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Access DB# 73098

# SEARCH REQUEST FORM

Scientific and Technical Information Center

Requester's Full Name: Dwayne C. Jones Examiner #: 71299 Date: 12 AUG 02  
Art Unit: 1614 Phone Number 301-4634 Serial Number: 89,899,683  
Mail Box and Bldg/Room Location: 2007, CM1 Results Format Preferred (circle) PAPER DISK E-MAIL

If more than one search is submitted, please prioritize searches in order of need.

\*\*\*\*\*  
Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: see attached sheet

Inventors (please provide full names): 11

Earliest Priority Filing Date: 13 JUL 2000

\*For Sequence Searches Only\* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

Please search <sup>method</sup> claims 1, 3, 11

and the composition claim of 16  
wherein the compounds of claim 3  
are used just as pharmaceuticals

Point of Contact:  
Barb O'Bryen  
Technical Information Specialist  
STIC CM1 6A05 308-4291

Point of Contact:  
Barb O'Bryen  
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Searcher Phone #: \_\_\_\_\_  
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Date Completed: 8-20-02  
Searcher Prep & Review Time: 50  
Clerical Prep Time: \_\_\_\_\_  
Online Time: 67

## Type of Search

NA Sequence (#) \_\_\_\_\_  
AA Sequence (#) \_\_\_\_\_  
Structure (#) \_\_\_\_\_  
Bibliographic X  
Litigation \_\_\_\_\_  
Fulltext \_\_\_\_\_  
Patent Family \_\_\_\_\_  
Other \_\_\_\_\_

## Vendors and cost where applicable

STN 299  
Dialog \_\_\_\_\_  
Questel/Orbit \_\_\_\_\_  
Dr.Link \_\_\_\_\_  
Lexis/Nexis \_\_\_\_\_  
Sequence Systems \_\_\_\_\_  
WWW/Internet \_\_\_\_\_  
Other (specify) \_\_\_\_\_

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epidermal edge of a sinus draining underlying osteomyelitis. **Meleney u.** undermining u. of the skin and subcutaneous tissues caused by a synergistic infection by microaerophilic nonhemolytic streptococci and aerobic hemolytic staphylococci. SYN: Meleney gangrene, progressive bacterial synergistic gangrene. **Mooren u.** chronic inflammation of the peripheral cornea that slowly progresses centrally with corneal thinning and sometimes perforation. **Oriental u.** the lesion occurring in cutaneous leishmaniasis. **penetrating u.** an u. extending into deeper tissues of an organ. **peptic u.** an u. of the alimentary mucosa, usually in the stomach or duodenum, exposed to acid gastric secretion. **perforated u.** an u. extending through the wall of an organ. **perforating u. of foot** a round, deep, trophic u. of the sole of the foot, following disease or injury, in any part of its course from the center to the periphery of the nerve supplying the part. **phagedenic u.** a rapidly spreading u. attended by the formation of extensive sloughing. SYN: sloughing u.. **phlegmonous u.** a u. accompanied by inflammation of the neighboring tissues. **pressure u.** SYN: decubitus u.. **recurrent aphthous u.** SYN: aphtha (2). **ring u. of cornea** inflammation of the greater part or the whole of the corneal periphery. **rodent u.** historic term for a slowly enlarging ulcerated basal cell carcinoma, usually on the face. **Saemisch u.** a form of serpiginous keratitis, frequently accompanied by hypopyon. **serpent u. of cornea** SYN: serpiginous keratitis. **serpiginous u.** an u. extending on one side while healing at the opposite edge, forming an undulating margin. **serpiginous corneal u.** serpentine ulceration of the cornea, due to infection, most often with Streptococcus pneumoniae. **simple u.** a local, not constitutional, u. not accompanied by marked pain or inflammation. **sloughing u.** SYN: phagedenic u.. **soft u.** SYN: chancroid. **stasis u.** SYN: varicose u.. **stercoral u.** an u. of the colon due to pressure and irritation of retained fecal masses. **stomal u.** an intestinal u. occurring after gastrojejunostomy in the jejunal mucosa near the opening (stoma) between the stomach and the jejunum. **Curling u.** SYN: stress u.. **Sutton u.** a solitary, deep, painful u. of the buccal or genital mucous membrane. **syphilitic u.** 1. SYN: chancre. 2. any ulceration caused by a syphilitic infection. **Syriac u.**, **Syrian u.** old names for diphtheria. **tanner's u.** SYN: chrome u.. **trophic u.** u. resulting from cutaneous sensory denervation. SEE ALSO: perforating u. of foot. SYN: trophic gangrene. **tropical u.** 1. the lesion occurring in cutaneous leishmaniasis; SYN: tropical sore. SEE ALSO: cutaneous leishmaniasis. 2. tropical phagedenic ulceration caused by a variety of microorganisms, including mycobacteria; common in northern Nigeria. **undermining u.** a chronic cutaneous u. with overhanging margins; due to hemolytic streptococci, tubercle bacilli, or other bacteria. **varicose u.** the loss of skin surface in the drainage area of a varicose vein, usually in the leg, resulting from stasis and infection. SEE ALSO: gravitational u.. SYN: stasis u., venous u.. **venereal u.** SYN: chancroid. **venous u.** SYN: varicose u.. **Zambesi u.** an u., usually single, about 3 cm in diameter, on the foot or leg, occurring in laborers in the Zambesi Delta; it has a sloughing surface, but does not spread and produces no constitutional symptoms or glandular enlargement; it is associated with the presence of a spirillum and a large fusiform bacillus; one attack seems to confer a partial immunity.

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STRUCTURE FILE UPDATES: 19 AUG 2002 HIGHEST RN 444278-83-5  
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TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when  
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Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES  
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Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> d rn cn 12

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 212126-32-4 REGISTRY  
CN 2-Cyclopenten-1-one, 2-(3,5-difluorophenyl)-3-[4-(methylsulfonyl)phenyl]-  
(9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN L 776967

=> d rn cn 13

L3 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 266320-83-6 REGISTRY  
CN 3(2H)-Pyridazinone, 2-(3,4-difluorophenyl)-4-(3-hydroxy-3-methylbutoxy)-5-  
[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)

=> d rn cn 14

L4 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 169590-42-5 REGISTRY  
CN Benzenesulfonamide, 4-[5-(4-methylphenyl)-3-(trifluoromethyl)-1H-pyrazol-1-  
yl]- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN 4-[5-(4-Methylphenyl)-3-(trifluoromethyl)-1H-pyrazol-1-  
yl]benzenesulfonamide  
CN Celebrex  
CN Celecoxib  
CN Celocoxib  
CN SC 58635  
CN YM 177

=> d rn cn 15

L5 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 169590-41-4 REGISTRY  
CN Benzenesulfonamide, 4-[3-(difluoromethyl)-5-(3-fluoro-4-methoxyphenyl)-1H-  
pyrazol-1-yl]- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN 4-[5-(3-Fluoro-4-methoxyphenyl)-3-(difluoromethyl)-1H-pyrazol-1-

yl]benzenesulfonamide

CN **Deracoxib**  
CN SC 046  
CN SC 46  
CN SC 59046

=> d rn cn 16

L6 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 181695-72-7 REGISTRY  
CN Benzenesulfonamide, 4-(5-methyl-3-phenyl-4-isoxazolyl)- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 4-(5-Methyl-3-phenylisoxazol-4-yl)benzenesulfonamide  
CN SC 65872  
CN **Valdecoxib**

=> d rn cn 17

L7 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 162011-90-7 REGISTRY  
CN 2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-phenyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 3-Phenyl-4-[4-(Methylsulfonyl)phenyl]-2(5H)-furanone  
CN MK 0966  
CN MK 966  
CN **Rofecoxib**  
CN Vioxx

=> d rn cn 18

L8 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 202409-33-4 REGISTRY  
CN 2,3'-Bipyridine, 5-chloro-6'-methyl-3-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 5-Chloro-6'-methyl-3-[4-(methylsulfonyl)phenyl]-2,3'-bipyridine  
CN **Etoricoxib**  
CN MK 0663  
CN MK 663

=> d rn cn

L10 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS  
RN 329900-75-6 REGISTRY  
CN Synthetase, prostaglandin endoperoxide, 2 (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Arachidonate cyclooxygenase 2  
CN COX 2  
CN **Cyclooxygenase 2**  
CN Prostaglandin endoperoxide H synthase-2  
CN Prostaglandin endoperoxide synthase-2  
CN Prostaglandin endoperoxide synthetase 2  
CN Prostaglandin G/H synthase-2  
CN Prostaglandin H synthase-2

=> fil cap1



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FILE COVERS 1907 - 20 Aug 2002 VOL 137 ISS 8  
FILE LAST UPDATED: 19 Aug 2002 (20020819/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> d que l18; d que l19; d que l21; d que l27; d que l31; d que l32; d que l33;d que l36

L9 1613 SEA FILE=CAPLUS ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE) (W) 2 (L) INHIBIT?/OBI  
L10 1 SEA FILE=REGISTRY ABB=ON "CYCLOOXYGENASE 2"/CN  
L11 532 SEA FILE=CAPLUS ABB=ON L10 (L) INHIBIT?/OBI  
L12 66 SEA FILE=CAPLUS ABB=ON PROSTAGLANDIN (2W) SYNTHASE (W) 2 (L) INHIBIT  
?/OBI  
L14 1245 SEA FILE=CAPLUS ABB=ON (BLEPHARITI? OR ENDOPHTHALMITI? OR  
EPISCLERITI? OR KERATITI? OR KERATOCONJUNCTIV? OR MOOREN?)/OBI  
L15 609 SEA FILE=CAPLUS ABB=ON CORNEA?/OBI (L) ?TRANSPLANT? OR RETINA? (L  
) DETACH?/OBI  
L16 370 SEA FILE=CAPLUS ABB=ON LENS## (L) (IMPLANT? OR ARTIFICIAL?)/OBI  
L18 1 SEA FILE=CAPLUS ABB=ON (L9 OR L11 OR L12) AND (L14 OR L15 OR  
L16)

L9 1613 SEA FILE=CAPLUS ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE) (W) 2 (L) INHIBIT?/OBI  
L10 1 SEA FILE=REGISTRY ABB=ON "CYCLOOXYGENASE 2"/CN  
L11 532 SEA FILE=CAPLUS ABB=ON L10 (L) INHIBIT?/OBI  
L12 66 SEA FILE=CAPLUS ABB=ON PROSTAGLANDIN (2W) SYNTHASE (W) 2 (L) INHIBIT  
?/OBI  
L13 14271 SEA FILE=CAPLUS ABB=ON EYE (L) (DISEASE# OR DISORDER#)/OBI  
L19 6 SEA FILE=CAPLUS ABB=ON (L9 OR L11 OR L12) (L) L13

L9 1613 SEA FILE=CAPLUS ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE) (W) 2 (L) INHIBIT?/OBI  
L10 1 SEA FILE=REGISTRY ABB=ON "CYCLOOXYGENASE 2"/CN  
L11 532 SEA FILE=CAPLUS ABB=ON L10 (L) INHIBIT?/OBI  
L12 66 SEA FILE=CAPLUS ABB=ON PROSTAGLANDIN (2W) SYNTHASE (W) 2 (L) INHIBIT  
?/OBI

L13 14271 SEA FILE=CAPLUS ABB=ON EYE(L) (DISEASE# OR DISORDER#)/OBI  
L20 6365 SEA FILE=CAPLUS ABB=ON OPHTHALM?/OBI  
L21 1 SEA FILE=CAPLUS ABB=ON (L9 OR L11 OR L12) AND L13 AND L20

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L14 1245 SEA FILE=CAPLUS ABB=ON (BLEPHARITI? OR ENDOPHTHALMITI? OR EPISCLERITI? OR KERATITI? OR KERATOCONJUNCTIV? OR MOOREN?)/OBI  
L15 609 SEA FILE=CAPLUS ABB=ON CORNEA?/OBI (L) ?TRANSPLANT? OR RETINA?(L) DETACH?/OBI  
L16 370 SEA FILE=CAPLUS ABB=ON LENS##(L) (IMPLANT? OR ARTIFICIAL?)/OBI

L24 544 SEA FILE=CAPLUS ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L25 421 SEA FILE=CAPLUS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)/OBI  
L26 85 SEA FILE=CAPLUS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)/OBI  
L27 1 SEA FILE=CAPLUS ABB=ON (L24 OR L25 OR L26) AND (L14 OR L15 OR L16)

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L13 14271 SEA FILE=CAPLUS ABB=ON EYE(L) (DISEASE# OR DISORDER#)/OBI  
L20 6365 SEA FILE=CAPLUS ABB=ON OPHTHALM?/OBI  
L24 544 SEA FILE=CAPLUS ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L25 421 SEA FILE=CAPLUS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)/OBI  
L26 85 SEA FILE=CAPLUS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)/OBI  
L31 2 SEA FILE=CAPLUS ABB=ON (L24 OR L25 OR L26) AND L20 AND L13

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF

L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L13 14271 SEA FILE=CAPLUS ABB=ON EYE(L) (DISEASE# OR DISORDER#)/OBI  
L20 6365 SEA FILE=CAPLUS ABB=ON OPHTHALM?/OBI  
L24 544 SEA FILE=CAPLUS ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L25 421 SEA FILE=CAPLUS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)/OBI  
L26 85 SEA FILE=CAPLUS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)/OBI  
L32 2 SEA FILE=CAPLUS ABB=ON (L24 OR L25 OR L26) (L) (L20 OR L13)

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L13 14271 SEA FILE=CAPLUS ABB=ON EYE(L) (DISEASE# OR DISORDER#)/OBI  
L22 4043 SEA FILE=CAPLUS ABB=ON L13(L) (PREVENT? OR TREAT? OR THERAP?)  
L24 544 SEA FILE=CAPLUS ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L25 421 SEA FILE=CAPLUS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)/OBI  
L26 85 SEA FILE=CAPLUS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)/OBI  
L33 7 SEA FILE=CAPLUS ABB=ON L22 AND (L24 OR L25 OR L26)

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L9 1613 SEA FILE=CAPLUS ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO OXYGENASE) (W) 2 (L) INHIBIT?/OBI  
L10 1 SEA FILE=REGISTRY ABB=ON "CYCLOOXYGENASE 2"/CN  
L11 532 SEA FILE=CAPLUS ABB=ON L10(L) INHIBIT?/OBI  
L12 66 SEA FILE=CAPLUS ABB=ON PROSTAGLANDIN(2W) SYNTHASE(W) 2 (L) INHIBIT ?/OBI  
L24 544 SEA FILE=CAPLUS ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L25 421 SEA FILE=CAPLUS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)/OBI  
L26 85 SEA FILE=CAPLUS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)/OBI

L35

226 SEA FILE=CAPLUS ABB=ON CORNEA?(L)INFLAM?/OBI

L36

0 SEA FILE=CAPLUS ABB=ON (L9 OR L11 OR L12 OR (L24 OR L25 OR L26)) AND L35

=&gt; s l18 or l19 or l21 or l27 or l31 or l32 or l33

L132

12 L18 OR L19 OR L21 OR L27 OR L31 OR L32 OR L33

=&gt; fil medl

FILE 'MEDLINE' ENTERED AT 16:52:04 ON 20 AUG 2002

FILE LAST UPDATED: 17 AUG 2002 (20020817/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2002 vocabulary. Enter HELP THESAURUS for details.

THIS FILE CONTAINS CAS REGISTRY NUMBERS FOR EASY AND ACCURATE SUBSTANCE IDENTIFICATION.

=&gt; d que 154; d que 156; d que 160; d que 173; d que 174; d que 189

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)

L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF

L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF

L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN

L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN

L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN

L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN

L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN

L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)

L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W)(046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)

L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W)(966 OR 0966) OR L 791456)

L41 646 SEA FILE=MEDLINE ABB=ON BLEPHARITIS/CT

L42 7272 SEA FILE=MEDLINE ABB=ON CORNEA+NT/CT(L)TR/CT OR CORNEAL TRANSPLANTATION+NT/CT

L43 2812 SEA FILE=MEDLINE ABB=ON ENDOPHTHALMITIS/CT

L44 384 SEA FILE=MEDLINE ABB=ON SCLERITIS/CT

L45 10979 SEA FILE=MEDLINE ABB=ON KERATITIS+NT/CT

L46 2397 SEA FILE=MEDLINE ABB=ON KERATOCONJUNCTIVITIS+NT/CT

L47 10861 SEA FILE=MEDLINE ABB=ON RETINAL DETACHMENT/CT

L48 5509 SEA FILE=MEDLINE ABB=ON LENS##(3A)(ARTIFICIAL OR IMPLANT?)

L49 157 SEA FILE=MEDLINE ABB=ON MOOREN?

L50 22221 SEA FILE=MEDLINE ABB=ON CORNEAL DISEASES+NT/CT

L51 2683 SEA FILE=MEDLINE ABB=ON CORNEAL ULCER/CT

L52 1755 SEA FILE=MEDLINE ABB=ON LENS IMPLANTATION, INTRAOCULAR/CT

L54 0 SEA FILE=MEDLINE ABB=ON (L38 OR L39 OR L40) AND ((L41 OR L42 OR L43 OR L44 OR L45 OR L46 OR L47 OR L48 OR L49 OR L50 OR L51 OR L52))

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR 169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33

-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR 329900-75-6/BI)

L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC 65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)  
L55 250466 SEA FILE=MEDLINE ABB=ON C11./CT *>eye disease*  
L56 4 SEA FILE=MEDLINE ABB=ON (L38 OR L39 OR L40) AND L55

L37 64351 SEA FILE=MEDLINE ABB=ON CYCLOOXYGENASE INHIBITORS+NT/CT  
L41 646 SEA FILE=MEDLINE ABB=ON BLEPHARITIS/CT  
L42 7272 SEA FILE=MEDLINE ABB=ON CORNEA+NT/CT (L)TR/CT OR CORNEAL TRANSPLANTATION+NT/CT  
L43 2812 SEA FILE=MEDLINE ABB=ON ENDOPHTHALMITIS/CT  
L44 384 SEA FILE=MEDLINE ABB=ON SCLERITIS/CT  
L45 10979 SEA FILE=MEDLINE ABB=ON KERATITIS+NT/CT  
L46 2397 SEA FILE=MEDLINE ABB=ON KERATOCONJUNCTIVITIS+NT/CT  
L47 10861 SEA FILE=MEDLINE ABB=ON RETINAL DETACHMENT/CT  
L48 5509 SEA FILE=MEDLINE ABB=ON LENS## (3A) (ARTIFICIAL OR IMPLANT?)  
L49 157 SEA FILE=MEDLINE ABB=ON MOOREN?  
L50 22221 SEA FILE=MEDLINE ABB=ON CORNEAL DISEASES+NT/CT  
L51 2683 SEA FILE=MEDLINE ABB=ON CORNEAL ULCER/CT  
L52 1755 SEA FILE=MEDLINE ABB=ON LENS IMPLANTATION, INTRAOCULAR/CT  
L58 5167 SEA FILE=MEDLINE ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO OXYGENASE OR PROSTAGLANDIN (2W)SYNTHASE) (W) 2  
L60 3 SEA FILE=MEDLINE ABB=ON L37 AND L58 AND (L41 OR L42 OR L43 OR L44 OR L45 OR L46 OR L47 OR L48 OR L49 OR L50 OR L51 OR L52)

L37 64351 SEA FILE=MEDLINE ABB=ON CYCLOOXYGENASE INHIBITORS+NT/CT  
L41 646 SEA FILE=MEDLINE ABB=ON BLEPHARITIS/CT  
L46 2397 SEA FILE=MEDLINE ABB=ON KERATOCONJUNCTIVITIS+NT/CT  
L49 157 SEA FILE=MEDLINE ABB=ON MOOREN?  
L73 3 SEA FILE=MEDLINE ABB=ON L37 AND (L41 OR L46 OR L49)

L37 64351 SEA FILE=MEDLINE ABB=ON CYCLOOXYGENASE INHIBITORS+NT/CT  
L51 2683 SEA FILE=MEDLINE ABB=ON CORNEAL ULCER/CT  
L74 6 SEA FILE=MEDLINE ABB=ON L37/MAJ AND L51/MAJ

L37 64351 SEA FILE=MEDLINE ABB=ON CYCLOOXYGENASE INHIBITORS+NT/CT  
L42 7272 SEA FILE=MEDLINE ABB=ON CORNEA+NT/CT (L)TR/CT OR CORNEAL TRANSPLANTATION+NT/CT  
L47 10861 SEA FILE=MEDLINE ABB=ON RETINAL DETACHMENT/CT  
L52 1755 SEA FILE=MEDLINE ABB=ON LENS IMPLANTATION, INTRAOCULAR/CT  
L86 11889 SEA FILE=MEDLINE ABB=ON PAIN, POSTOPERATIVE/CT  
L89 3 SEA FILE=MEDLINE ABB=ON (L42 OR L47 OR L52) AND L86 AND L37

=> s 156 or 160 or 173 or 174 or 189

L133 18 L56 OR L60 OR L73 OR L74 OR L89

=> fil wpids

FILE 'WPIDS' ENTERED AT 16:52:07 ON 20 AUG 2002  
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FILE LAST UPDATED: 15 AUG 2002 <20020815/UP>  
MOST RECENT DERWENT UPDATE 200252 <200252/DW>  
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> The BATCH option for structure searches has been  
enabled in WPINDEX/WPIDS and WPIX >>>

>>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY >>>

>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES,  
SEE <http://www.derwent.com/dwpi/updates/dwpcov/index.html> <<<

>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,  
PLEASE VISIT:  
[http://www.stn-international.de/training\\_center/patents/stn\\_guide.pdf](http://www.stn-international.de/training_center/patents/stn_guide.pdf) <<<

>>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER  
GUIDES, PLEASE VISIT:  
[http://www.derwent.com/userguides/dwpi\\_guide.html](http://www.derwent.com/userguides/dwpi_guide.html) <<<

=> d que 199; d que 1100; d que 1103

L92 442 SEA FILE=WPIDS ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE OR PROSTAGLANDIN(2W)SYNTHASE) (W)2(3A)INHIBIT?  
L93 22 SEA FILE=WPIDS ABB=ON COX2 (3A)INHIBIT?  
L96 743 SEA FILE=WPIDS ABB=ON BLEPHARITI? OR ENDOPHTHALMITI? OR  
EPISCLERITI? OR KERATITI? OR KERATOCONJUNCTIV? OR MOOREN?  
L97 2090 SEA FILE=WPIDS ABB=ON RETINA?(2A)DETACH? OR LENS##(3A)(IMPLANT  
? OR ARTIFICIAL?) OR CORNEA?(3A)?TRANSPLANT?  
L98 641 SEA FILE=WPIDS ABB=ON CORNEA?(3A)(INFLAM? OR ULCER?)  
L99 5 SEA FILE=WPIDS ABB=ON (L92 OR L93) AND (L96 OR L97 OR L98)

L94 92 SEA FILE=WPIDS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR  
SC(W)(046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L95 23 SEA FILE=WPIDS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W)(966 OR 0966) OR L 791456)  
L96 743 SEA FILE=WPIDS ABB=ON BLEPHARITI? OR ENDOPHTHALMITI? OR  
EPISCLERITI? OR KERATITI? OR KERATOCONJUNCTIV? OR MOOREN?  
L97 2090 SEA FILE=WPIDS ABB=ON RETINA?(2A)DETACH? OR LENS##(3A)(IMPLANT  
? OR ARTIFICIAL?) OR CORNEA?(3A)?TRANSPLANT?  
L98 641 SEA FILE=WPIDS ABB=ON CORNEA?(3A)(INFLAM? OR ULCER?)  
L100 2 SEA FILE=WPIDS ABB=ON (L94 OR L95) AND (L96 OR L97 OR L98)

L94 92 SEA FILE=WPIDS ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB OR  
SC(W)(046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L95 23 SEA FILE=WPIDS ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W)(966 OR 0966) OR L 791456)  
L101 10277 SEA FILE=WPIDS ABB=ON OPHTHALM?  
L103 4 SEA FILE=WPIDS ABB=ON L101 (S) (L94 OR L95)

=> s 199 or 1100 or 1103

L134 9 L99 OR L100 OR L103

=> fil embase

FILE 'EMBASE' ENTERED AT 16:52:11 ON 20 AUG 2002  
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FILE COVERS 1974 TO 15 Aug 2002 (20020815/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate  
substance identification.

=> d que 1116; d que 1120

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR  
169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33  
-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR  
329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR  
L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB  
OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)  
L104 2385 SEA FILE=EMBASE ABB=ON (L38 OR L39 OR L40)  
L105 2504 SEA FILE=EMBASE ABB=ON CYCLOOXYGENASE 2 INHIBITOR/CT OR COX 2  
INHIBITOR/CT  
L106 665 SEA FILE=EMBASE ABB=ON BLEPHARITIS/CT  
L107 3720 SEA FILE=EMBASE ABB=ON CORNEA TRANSPLANTATION/CT  
L108 2799 SEA FILE=EMBASE ABB=ON ENDOPHTHALMITIS/CT  
L109 673 SEA FILE=EMBASE ABB=ON SCLERITIS/CT  
L110 4769 SEA FILE=EMBASE ABB=ON KERATITIS/CT  
L111 2021 SEA FILE=EMBASE ABB=ON KERATOCONJUNCTIVITIS/CT OR KERATOCONJUN  
CTIVITIS SICCA/CT  
L112 7695 SEA FILE=EMBASE ABB=ON RETINA DETACHMENT/CT  
L113 723 SEA FILE=EMBASE ABB=ON LENS IMPLANTATION/CT  
L114 145 SEA FILE=EMBASE ABB=ON CORNEA RODENT ULCER/CT  
L115 140 SEA FILE=EMBASE ABB=ON MOOREN?  
L116 2 SEA FILE=EMBASE ABB=ON (L104 OR L105) AND (L106 OR L107 OR  
L108 OR L109 OR L110 OR L111 OR L112 OR L113 OR L114 OR L115)

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR  
169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33  
-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR  
329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN

L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR  
L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CELCOXIB OR DERACOXIB  
OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)  
L104 2385 SEA FILE=EMBASE ABB=ON (L38 OR L39 OR L40)  
L105 2504 SEA FILE=EMBASE ABB=ON CYCLOOXYGENASE 2 INHIBITOR/CT OR COX 2  
INHIBITOR/CT  
L119 2473 SEA FILE=EMBASE ABB=ON EYE DROPS/CT  
L120 5 SEA FILE=EMBASE ABB=ON (L104 OR L105) AND L119

=> s 1116 or 1120

L135 5 L116 OR L120

=> fil drugu

FILE 'DRUGU' ENTERED AT 16:52:14 ON 20 AUG 2002  
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FILE LAST UPDATED: 15 AUG 2002 <20020815/UP>  
>>> DERWENT DRUG FILE (SUBSCRIBER) <<<

>>> SDI'S MAY BE RUN WEEKLY OR MONTHLY AS OF JUNE 2001. <<<  
>>> (WEEKLY IS THE DEFAULT). FOR PRICING INFORMATION <<<  
>>> SEE HELP COST <<<

>>> FILE COVERS 1983 TO DATE <<<  
>>> THESAURUS AVAILABLE IN /CT <<<

=> d que 1127;d que 1129;d que 1131

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR  
169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33  
-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR  
329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR  
L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CELCOXIB OR DERACOXIB  
OR SC(W) (046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W) (966 OR 0966) OR L 791456)  
L121 885 SEA FILE=DRUGU ABB=ON (L38 OR L39 OR L40)  
L122 3134 SEA FILE=DRUGU ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE OR PROSTAGLANDIN(2W)SYNTHASE) (W) 2(3A) INHIBIT?  
L123 40 SEA FILE=DRUGU ABB=ON COX2(3A)INHIBIT?  
L124 1375 SEA FILE=DRUGU ABB=ON BLEPHARITI? OR ENDOPHTHALMITI? OR  
EPISCLERITI? OR KERATITI? OR KERATOCONJUNCTIV? OR MOOREN?  
L125 439 SEA FILE=DRUGU ABB=ON RETINA?(2A)DETACH? OR LENS##(3A) (IMPLANT



? OR ARTIFICIAL?) OR CORNEA?(3A)?TRANSPLANT?  
L126 278 SEA FILE=DRUGU ABB=ON CORNEA?(2A)(ULCER? OR INFLAMM?)  
L127 2 SEA FILE=DRUGU ABB=ON (L121 OR L122 OR L123) AND (L124 OR  
L125 OR L126)  
  
L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR  
169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33  
-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR  
329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR  
L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB  
OR SC(W)(046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W)(966 OR 0966) OR L 791456)  
L121 885 SEA FILE=DRUGU ABB=ON (L38 OR L39 OR L40)  
L122 3134 SEA FILE=DRUGU ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE OR PROSTAGLANDIN(2W)SYNTHASE)(W)2(3A)INHIBIT?  
L123 40 SEA FILE=DRUGU ABB=ON COX2(3A)INHIBIT?  
L128 3378 SEA FILE=DRUGU ABB=ON OPHTHALM?/CT  
L129 2 SEA FILE=DRUGU ABB=ON (L121 OR L122 OR L123) AND L128

L1 10 SEA FILE=REGISTRY ABB=ON (162011-90-7/BI OR 169590-41-4/BI OR  
169590-42-5/BI OR 181695-72-7/BI OR 198470-84-7/BI OR 202409-33  
-4/BI OR 212126-32-4/BI OR 254-04-6/BI OR 266320-83-6/BI OR  
329900-75-6/BI)  
L2 1 SEA FILE=REGISTRY ABB=ON L1 AND C18H14F2O3S/MF  
L3 1 SEA FILE=REGISTRY ABB=ON L1 AND C22H22F2N2O5S/MF  
L4 1 SEA FILE=REGISTRY ABB=ON CELECOXIB/CN  
L5 1 SEA FILE=REGISTRY ABB=ON DERACOXIB/CN  
L6 1 SEA FILE=REGISTRY ABB=ON VALDECOXIB/CN  
L7 1 SEA FILE=REGISTRY ABB=ON ROFECOXIB/CN  
L8 1 SEA FILE=REGISTRY ABB=ON ETORICOXIB/CN  
L38 427 SEA FILE=MEDLINE ABB=ON (L2 OR L3 OR L4 OR L5 OR L6 OR L7 OR  
L8)  
L39 622 SEA FILE=MEDLINE ABB=ON (CELEBREX OR CEL!COXIB OR DERACOXIB  
OR SC(W)(046 OR 46 OR 59046) OR VALDECOXIB OR BEXTRA)  
L40 82 SEA FILE=MEDLINE ABB=ON (YM 177 OR SC 58635 OR VIOXX OR SC  
65872 OR ETORICOXIB OR MK(W)(966 OR 0966) OR L 791456)  
L121 885 SEA FILE=DRUGU ABB=ON (L38 OR L39 OR L40)  
L122 3134 SEA FILE=DRUGU ABB=ON (COX OR CYCLOOXYGENASE OR CYCLO  
OXYGENASE OR PROSTAGLANDIN(2W)SYNTHASE)(W)2(3A)INHIBIT?  
L123 40 SEA FILE=DRUGU ABB=ON COX2(3A)INHIBIT?  
L130 4763 SEA FILE=DRUGU ABB=ON OPHTHALMOLOGICAL/CC  
L131 6 SEA FILE=DRUGU ABB=ON (L121 OR L122 OR L123) AND L130

=> s 1127 or 1129 or 1131

L136 9 L127 OR L129 OR L131

=> dup rem 1133,1136,1135,1132,1134

FILE 'MEDLINE' ENTERED AT 16:53:10 ON 20 AUG 2002

FILE 'DRUGU' ENTERED AT 16:53:10 ON 20 AUG 2002  
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PROCESSING COMPLETED FOR L133

PROCESSING COMPLETED FOR L136

PROCESSING COMPLETED FOR L135

PROCESSING COMPLETED FOR L132

PROCESSING COMPLETED FOR L134

L137

50 DUP REM L133 L136 L135 L132 L134 (3 DUPLICATES REMOVED)

ANSWERS '1-18' FROM FILE MEDLINE

ANSWERS '19-27' FROM FILE DRUGU

ANSWERS '28-32' FROM FILE EMBASE

ANSWERS '33-44' FROM FILE CAPLUS

ANSWERS '45-50' FROM FILE WPIDS

=> d ibib ab hitrn 1-50; fil hom

L137 ANSWER (1 OF 50)

MEDLINE

ACCESSION NUMBER: 2002106815 MEDLINE

DOCUMENT NUMBER: 21679446 PubMed ID: 11821217

TITLE: Naproxen ophthalmic solution to manage inflammation after phacoemulsification.

AUTHOR: Papa Vincenzo; Milazzo Giovanni; Santocono Marcello; Servolle Valerie; Sourdille Philippe; Santiago Pierre-Yves; Darondeau Jacques; Cassoux Nathalie; LeHoang Phuc  
CORPORATE SOURCE: Medical Department SIFI S.p.A, Lavinaio-Catania, Italy..

SOURCE: JOURNAL OF CATARACT AND REFRACTIVE SURGERY, (2002 Feb) 28 (2) 321-7.

PUB. COUNTRY: Journal code: 8604171. ISSN: 0886-3350.

DOCUMENT TYPE: United States  
(CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)

(MULTICENTER STUDY)  
(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200203

ENTRY DATE: Entered STN: 20020213

Last Updated on STN: 20020312

Entered Medline: 20020311

AB PURPOSE: To explore the efficacy and safety of 2 concentrations (0.1% and 0.2%) of sodium naproxen ophthalmic solution in controlling ocular inflammation in patients having phacoemulsification and intraocular lens implantation. SETTING: Service d'Ophthalmologie La Pitie' and Centre Ophthalmologique, Paris, and Clinique Sourdille, Nantes, France; Department of Ophthalmology, University of Lausanne, Switzerland. METHODS: One hundred one patients were randomly treated with naproxen 0.1%, naproxen 0.2%, or diclofenac 0.1% 3 times a day for 30 days starting the day before surgery. Postsurgical ocular inflammation was measured after 1, 10, and 30 days using the Kowa FC-1000 laser flare-cell meter and a conventional

slitlamp biomicroscope. Safety parameters were evaluated at the same visits. RESULTS: Naproxen 0.2% ophthalmic solution and diclofenac 0.01% were comparable in controlling postsurgical inflammation. The naproxen was well tolerated. No serious adverse events occurred during the study. CONCLUSIONS: These preliminary results suggest that naproxen ophthalmic solution may be effectively and safely used to control inflammation after uneventful phacoemulsification. Because of the limited number of patients, larger studies are needed to confirm these results.

L137 ANSWER 2 OF 50 MEDLINE  
ACCESSION NUMBER: 2001672527 MEDLINE  
DOCUMENT NUMBER: 21575000 PubMed ID: 11718490  
TITLE: Visual disturbance associated with celecoxib--a  
comment.  
COMMENT: Comment on: Pharmacotherapy. 2001 Jan;21(1):114-5  
AUTHOR: Gehrs K M  
CORPORATE SOURCE: Department of Ophthalmology, University of Iowa Hospitals  
and Clinics, Iowa City 52242, USA.  
SOURCE: PHARMACOTHERAPY, (2001 Aug) 21 (8) 1014.  
Journal code: 8111305. ISSN: 0277-0008.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Commentary  
Letter  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200205  
ENTRY DATE: Entered STN: 20011126  
Last Updated on STN: 20020522  
Entered Medline: 20020517

L137 ANSWER 3 OF 50 MEDLINE  
ACCESSION NUMBER: 2001231537 MEDLINE  
DOCUMENT NUMBER: 21220975 PubMed ID: 11320025  
TITLE: Keratitis, ulceration, and perforation associated with  
topical nonsteroidal anti-inflammatory drugs.  
AUTHOR: Guidera A C; Luchs J I; Udell I J  
CORPORATE SOURCE: Department of Ophthalmology, Long Island Jewish Medical  
Center, New Hyde Park, New York, NY, USA.  
SOURCE: OPHTHALMOLOGY, (2001 May) 108 (5) 936-44.  
Journal code: 7802443. ISSN: 0161-6420.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200105  
ENTRY DATE: Entered STN: 20010517  
Last Updated on STN: 20010517  
Entered Medline: 20010510

AB PURPOSE: To report corneal complications associated with topical  
nonsteroidal anti-inflammatory drugs (NSAIDs). DESIGN: Retrospective,  
noncomparative interventional case series. PARTICIPANTS: Eighteen eyes of  
16 patients with adverse corneal events associated with NSAID use.  
METHODS: Evaluation of 16 patients referred for management of corneal  
complications during use of topical NSAIDs (ketorolac tromethamine  
[Acular], diclofenac sodium [Voltaren], diclofenac sodium [Falcon DSOS]).  
MAIN OUTCOME MEASURES: Type and severity of corneal complications.  
RESULTS: Of the 16 patients, two experienced severe keratopathy, three  
experienced ulceration, six experienced corneal or scleral melts, and five  
experienced perforations. Eleven patients had recent cataract surgery;  
nine of these were on concurrent topical steroids and antibiotics. Another  
patient who did not have recent surgery was using concurrent topical  
steroids without antibiotics for sarcoid uveitis. Systemic associations  
included two patients with rheumatoid arthritis, one patient with

asymptomatic Sjogren's syndrome, and two with rosacea. CONCLUSIONS: Topical NSAIDs were associated with corneal complications in 18 eyes of 16 patients. Potential risk factors include conditions that predispose the patient to corneal melting, concurrent topical steroids, and epithelial keratopathy in the early postoperative period.

L137 ANSWER 4 OF 50

MEDLINE

ACCESSION NUMBER:

2001208758

MEDLINE

DOCUMENT NUMBER:

21196009

PubMed ID: 11297478

TITLE:

The role of matrix metalloproteinases in ulcerative keratolysis associated with perioperative diclofenac use.  
O'Brien T P; Li Q J; Sauerburger F; Reviglio V E; Rana T; Ashraf M F

AUTHOR:

CORPORATE SOURCE:

Ocular Microbiology and Immunology Laboratory, The Wilmer Eye Institute, Johns Hopkins University School of Medicine, 600 N. Wolfe Street, Woods Bldg./Rm. 259, Baltimore, MD 21287-9121, USA.

SOURCE:

OPHTHALMOLOGY, (2001 Apr) 108 (4) 656-9.

Journal code: 7802443. ISSN: 0161-6420.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200104

ENTRY DATE:

Entered STN: 20010425

Last Updated on STN: 20010425

Entered Medline: 20010419

AB

OBJECTIVE: To investigate the role of matrix metalloproteinases (MMPs) in the pathogenesis of ulcerative keratolysis associated with topical use of generic diclofenac preoperatively and postoperatively. To characterize the inflammatory response of the cornea in this case of ulcerative keratolysis. DESIGN: Case report with clinicopathologic correlation. MAIN OUTCOME MEASURES: Corneal culture for microbial growth. Clinical and histopathologic examinations including routine histopathologic, immunofluorescent, and immunohistochemical studies. RESULTS: Microscopic examination of the corneal button disclosed fibrinous material with neutrophils and mononuclear inflammatory cells. The corneal epithelial basement membrane was irregularly thickened and patchy. Immunohistochemical staining detected weak staining of MMP-1 and a strong presence of MMP-8 in the epithelium. MMP-8 and 9 were also present in areas of leukocytic infiltration. MMP-2 appeared in a few stromal cells. Macrophages and leukocytes were the predominant infiltrating cells. CONCLUSIONS: A nonspecific inflammatory response occurred in this case of ulcerative keratolysis. Corneal epithelial cells are capable of secreting MMP-1 and 8 and may participate in the stromal degradation and repair process of the ulcerative keratolysis associated with topical nonsteroidol antiinflammatory use.

L137 ANSWER 5 OF 50

MEDLINE

ACCESSION NUMBER:

2001175672

MEDLINE

DOCUMENT NUMBER:

21170593

PubMed ID: 11270263

TITLE:

[64th Congress of the American College of Rheumatology, Philadelphia, October 28-November 2, 2000].  
64e congres de l'American College of Rheumatology, Philadelphia, 28 octobre-2 novembre 2000.

AUTHOR:

Hachulla E

CORPORATE SOURCE:

Service de medecine interne, hopital Claude-Huriez, place de Verdun, 59037 Lille, France.

SOURCE:

REVUE DE MEDECINE INTERNE, (2001 Mar) 22 (3) 219-27.

Journal code: 8101383. ISSN: 0248-8663.

PUB. COUNTRY:

France

DOCUMENT TYPE:

Conference; Conference Article; (CONGRESSES)

LANGUAGE:

French

FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200106  
ENTRY DATE: Entered STN: 20010611  
Last Updated on STN: 20010611  
Entered Medline: 20010607

L137 ANSWER 6 OF 50 MEDLINE  
ACCESSION NUMBER: 2001146245 MEDLINE  
DOCUMENT NUMBER: 21030579 PubMed ID: 11191731  
TITLE: Visual disturbance associated with celecoxib.  
COMMENT: Comment in: Pharmacotherapy. 2001 Aug;21(8):1014  
AUTHOR: Lund B C; Neiman R F  
CORPORATE SOURCE: Clinical and Administrative Division, College of Pharmacy  
Iowa City, IA 52242-1112, USA.. brian-lund@uiowa.edu  
SOURCE: PHARMACOTHERAPY, (2001 Jan) 21 (1) 114-5.  
Journal code: 8111305. ISSN: 0277-0008.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200103  
ENTRY DATE: Entered STN: 20010404  
Last Updated on STN: 20020522  
Entered Medline: 20010315

AB Celecoxib, a specific inhibitor of cyclooxygenase-2, is used to treat the symptoms of arthritis. A 79-year-old woman developed an atypical visual disturbance associated with this agent that resolved on discontinuation of celecoxib. Similar visual disturbances described with the traditional nonsteroidal antiinflammatory drugs are discussed.

L137 ANSWER 7 OF 50 MEDLINE  
ACCESSION NUMBER: 2001017575 MEDLINE  
DOCUMENT NUMBER: 20476690 PubMed ID: 11020603  
TITLE: New pieces for the puzzle: nonsteroidal anti-inflammatory drugs and corneal ulcers.  
AUTHOR: Price F W  
SOURCE: JOURNAL OF CATARACT AND REFRACTIVE SURGERY, (2000 Sep) 26 (9) 1263-5. Ref: 15  
Journal code: 8604171. ISSN: 0886-3350.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Editorial  
General Review; (REVIEW)  
(REVIEW, TUTORIAL)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200011  
ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20001107

L137 ANSWER 8 OF 50 MEDLINE  
ACCESSION NUMBER: 2000037164 MEDLINE  
DOCUMENT NUMBER: 20037164 PubMed ID: 10570586  
TITLE: Rheumatoid arthritis.  
COMMENT: Comment on: J Am Dent Assoc. 1999 May;130(5):689-98  
AUTHOR: Rosenstein E D; Kushner L J; Kramer N  
SOURCE: JOURNAL OF THE AMERICAN DENTAL ASSOCIATION, (1999 Oct) 130 (10) 1424, 1426.  
Journal code: 7503060. ISSN: 0002-8177.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Commentary  
Letter

LANGUAGE: English  
FILE SEGMENT: Dental Journals; Priority Journals  
ENTRY MONTH: 199911  
ENTRY DATE: Entered STN: 20000111  
Last Updated on STN: 20000229  
Entered Medline: 19991123

L137 ANSWER 9 OF 50

MEDLINE

ACCESSION NUMBER: 1999451103 MEDLINE  
DOCUMENT NUMBER: 99451103 PubMed ID: 10520225  
TITLE: The effect of selective cyclooxygenase-2 inhibitor on corneal angiogenesis in the rat.  
AUTHOR: Yamada M; Kawai M; Kawai Y; Mashima Y  
CORPORATE SOURCE: Department of Ophthalmology, Keio University School of Medicine, Tokyo, Japan. yamadam@med.keio.ac.jp  
SOURCE: CURRENT EYE RESEARCH, (1999 Oct) 19 (4) 300-4.  
Journal code: 8104312. ISSN: 0271-3683.  
PUB. COUNTRY: ENGLAND: United Kingdom  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199912  
ENTRY DATE: Entered STN: 20000113  
Last Updated on STN: 20000113  
Entered Medline: 19991213

AB PURPOSE. Eicosanoids that are present in inflamed tissues are thought to play a significant role in angiogenesis. Cyclooxygenase, a key enzyme in eicosanoid synthesis, has recently been shown to exist in two isoforms: the constitutive COX-1 and the inducible COX-2. This study was undertaken to determine the role of COX-2 in the corneal angiogenic response. METHODS. Angiogenesis in the rat cornea was provoked by chemical cautery. Either NS-398, a selective COX-2 inhibitor, or indomethacin, a non-selective COX inhibitor, was applied topically 3 times daily for 4 days. Neovascularization was quantitated by digital image analysis in corneal flat preparations. To test their inhibitory effects on eicosanoid synthesis, normal or cauterized corneas were incubated in the culture medium with the inhibitor. Prostaglandin E2 in the medium was assayed using an enzyme-linked immunosorbent assay. RESULTS. Both NS-398 and indomethacin significantly inhibited corneal neovascularization with the % inhibition of 36.4 +/- 9.6%, and 38.5 +/- 9.0%, respectively, when applied topically at a concentration of 0.1% (p < .001). Neither reduced the angiogenic response at a concentration of 0.01% or below. PGE(2) production in the cauterized cornea was 2.0 times higher than that in the controls. In normal corneas, indomethacin inhibited PGE(2) synthesis by 80%, whereas NS-398 inhibited it by no more than 20%. In contrast, in injured corneas, both indomethacin and NS-398 inhibited PGE(2) synthesis in a similar fashion, with a maximal inhibition rate of 75 to 80%. CONCLUSIONS. Our results suggest that COX-2 induction in cauterized corneas increases the level of eicosanoids, which result in corneal angiogenesis.

L137 ANSWER 10 OF 50

MEDLINE

ACCESSION NUMBER: 1999186594 MEDLINE  
DOCUMENT NUMBER: 99186594 PubMed ID: 10088733  
TITLE: Ketorolac tromethamine 0.5% ophthalmic solution in the treatment of moderate to severe ocular inflammation after cataract surgery: a randomized, vehicle-controlled clinical trial.  
COMMENT: Comment in: Am J Ophthalmol. 1999 Nov;128(5):662-3  
AUTHOR: Heier J; Cheetham J K; Degryse R; Dirks M S; Caldwell D R; Silverstone D E; Rosenthal A  
CORPORATE SOURCE: Ophthalmic Consultants of Boston and Center for Eye

SOURCE: Research, Massachusetts, USA.  
AMERICAN JOURNAL OF OPHTHALMOLOGY, (1999-Mar) 127 (3)  
253-9.  
Journal code: 0370500. ISSN: 0002-9394.

PUB. COUNTRY: United States

DOCUMENT TYPE: (CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)  
(MULTICENTER STUDY)  
(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199903

ENTRY DATE: Entered STN: 19990402  
Last Updated on STN: 20000327  
Entered Medline: 19990324

AB PURPOSE: To investigate the efficacy and safety of ketorolac tromethamine 0.5% ophthalmic solution (Acular; Allergan, Inc, Irvine, California) in the treatment of moderate to severe anterior segment inflammation developing after unilateral cataract surgery with intraocular lens implantation. METHODS: Only patients who exhibited moderate or greater levels of cells and flare 1 day after surgery were included in this multicenter, double-masked, randomly assigned, parallel-group study. Topical ketorolac or vehicle solution (Allergan, Inc) was administered to the treated eye four times daily, starting the day after surgery and continuing for 14 days. RESULTS: Ketorolac was significantly more effective than the vehicle solution in reducing anterior chamber cells ( $P < \text{or} = .030$ ) and flare ( $P < \text{or} = .025$ ), conjunctival erythema ( $P < \text{or} = .046$ ), ciliary flush ( $P < \text{or} = .006$ ), tearing ( $P < \text{or} = .012$ ), photophobia ( $P < \text{or} = .014$ ), and pain ( $P < \text{or} = .049$ ). Half as many patients from the ketorolac group (14/51) were discontinued from the study for lack of efficacy, compared with the vehicle group (28/51;  $P = .005$ ). There was no significant difference between ketorolac and the vehicle solution in changes in visual acuity, intraocular pressure, biomicroscopic or ophthalmoscopic variables, or adverse events. CONCLUSIONS: Ketorolac, tromethamine 0.5% ophthalmic solution is safe and provides substantial anti-inflammatory activity in the treatment of moderate to severe anterior segment inflammation developing after cataract surgery and intraocular lens implantation.

L137 ANSWER 11 OF 50 MEDLINE

ACCESSION NUMBER: 1998347625 MEDLINE

DOCUMENT NUMBER: 98347625 PubMed ID: 9682703

TITLE: The effects of topical nonsteroidal anti-inflammatory drugs on adenoviral replication.

AUTHOR: Gordon Y J; Araullo-Cruz T; Romanowski E G

CORPORATE SOURCE: Department of Ophthalmology, University of Pittsburgh School of Medicine, Pa., USA.. yjgordon@vision.eei.upmc.edu

CONTRACT NUMBER: EY05232 (NEI)

SOURCE: ARCHIVES OF OPHTHALMOLOGY, (1998-Jul) 116 (7) 900-5.  
Journal code: 7706534. ISSN: 0003-9950.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199808

ENTRY DATE: Entered STN: 19980817  
Last Updated on STN: 20000303  
Entered Medline: 19980804

AB OBJECTIVE: To evaluate the antiviral activity of topical diclofenac (Voltaren Ophthalmic) and ketorolac tromethamine (Acular) (2 non anti-inflammatory drugs [NSAIDs]) on adenoviral replication in the adenovirus (Ad) 5 McEwen-New Zealand rabbit ocular model. The 50% inhibitory concentration of ketorolac and diclofenac and

respective preservative components were determined for common ocular adenoviral serotypes (Ad8, Ad19, Ad1, and Ad5). In a series of experiments, Ad5 McEwen-inoculated New Zealand rabbit eyes were treated topically 4 times daily for 18 days with either ketorolac, diclofenac, prednisolone acetate (Pred Forte), or control vehicle (Comfort Tears). MAIN OUTCOME MEASURES: Outcome measures included serial ocular tear film titers and the formation of subepithelial immune corneal infiltrates. RESULTS: In vitro, neither ketorolac nor diclofenac demonstrated significant inhibitory activity against Ad1, Ad5, Ad8, or Ad19. In the rabbit model, there were no statistically significant differences among ketorolac, diclofenac, and the control vehicle with respect to viral replication or the formation of subepithelial immune infiltrates. In contrast, 1% prednisolone prolonged viral shedding and inhibited immune infiltrates ( $P < .001$  for both). CONCLUSIONS: Our experimental study suggests that treatment of epidemic keratoconjunctivitis with topical NSAIDs may be a safer alternative than topical steroids. Only controlled clinical trials can determine whether topical NSAIDs can provide symptomatic relief and not interfere with normal viral clearance.

L137 ANSWER 12 OF 50

MEDLINE

ACCESSION NUMBER: 1998287247 MEDLINE

DOCUMENT NUMBER: 98287247 PubMed ID: 9625565

TITLE: Topical diclofenac sodium in the management of anesthetic abuse keratopathy.

AUTHOR: Dornic D I; Thomas J M; Lass J H

CORPORATE SOURCE: Department of Ophthalmology, University Hospitals of Cleveland, and Case Western Reserve University School of Medicine, OH 44106, USA.

SOURCE: AMERICAN JOURNAL OF OPHTHALMOLOGY, (1998 May) 125 (5) 719-21.

Journal code: 0370500. ISSN: 0002-9394.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199806

ENTRY DATE: Entered STN: 19980625

Last Updated on STN: 19980625

Entered Medline: 19980615

AB PURPOSE: To report a case of anesthetic abuse keratopathy and to suggest the use of topical diclofenac sodium in the management of this disorder. METHOD: Narcotics and topical diclofenac were used to control pain in a patient who developed a corneal ulcer after abusing topical anesthetics. RESULT: After the institution of topical diclofenac, the patient reported substantial improvement in comfort and less reliance on narcotic agents for analgesia. CONCLUSION: We found topical diclofenac to be useful in controlling pain in this patient with anesthetic abuse keratopathy.

L137 ANSWER 13 OF 50

MEDLINE

ACCESSION NUMBER: 1998304377 MEDLINE

DOCUMENT NUMBER: 98304377 PubMed ID: 9640195

TITLE: Use of indomethacin for pain relief following scleral buckling surgery.

AUTHOR: Sadiq S A; Stevenson L; Gorman C; Orr G M

CORPORATE SOURCE: Department of Ophthalmology, Queen's Medical Centre, Nottingham.

SOURCE: BRITISH JOURNAL OF OPHTHALMOLOGY, (1998 Apr) 82 (4) 429-31. Journal code: 0421041. ISSN: 0007-1161.

COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

English



FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199807  
ENTRY DATE: Entered STN: 19980716  
Last Updated on STN: 19980716  
Entered Medline: 19980706

AB BACKGROUND/AIMS: Patients undergoing scleral buckling and cryotherapy suffer from mild to moderate postoperative pain. As good pain relief facilitates post-operative ocular examination, as well as patient comfort and recovery, the authors designed a prospective randomised double masked trial to evaluate the efficacy of indomethacin as a satisfactory analgesic for such patients. METHOD: Patients with a primary uncomplicated rhegmatogenous retinal detachment requiring scleral buckling and cryotherapy were randomly allocated to receive either indomethacin or placebo. A rectal suppository was administered 2 hours before surgery, followed by two capsules twice daily for 10 days. Pain relief was assessed with a linear graphic rating scale at the end of each day. Supplementary analgesia was allowed and recorded. RESULTS: 12 patients received indomethacin (group A) and 16 received placebo (group B). The extent of surgery was similar in both groups. One patient in group A, and two in group B withdrew after 3 days. The pain scores were converted to changes from the baseline (score on day 1), and the area under the curve calculated for each patient. The means of the areas were analysed with the Mann-Whitney test and showed that indomethacin caused a statistically significant reduction in pain score, both at 3 days ( $p = 0.04$ ) and at 10 days ( $p = 0.014$ ). There was no statistically significant difference in extra analgesic requirements between the two groups ( $p = 0.2$ ). CONCLUSIONS: Indomethacin is recommended for short to medium term pain relief following scleral buckling and cryotherapy.

L137 ANSWER 14 OF 50 MEDLINE  
ACCESSION NUMBER: 97298858 MEDLINE  
DOCUMENT NUMBER: 97298858 PubMed ID: 9154274  
TITLE: Cyclooxygenase-2 inhibitors: a new approach to the therapy of ocular inflammation.  
AUTHOR: Masferrer J L; Kulkarni P S  
CORPORATE SOURCE: G. D. Searle/Monsanto, St. Louis, Missouri, USA.  
SOURCE: SURVEY OF OPHTHALMOLOGY, 41 Suppl 2 S35-40.  
Ref: 43  
Journal code: 0404551. ISSN: 0039-6257.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
General Review; (REVIEW)  
(REVIEW, TUTORIAL)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199706  
ENTRY DATE: Entered STN: 19970620  
Last Updated on STN: 19970620  
Entered Medline: 19970612

AB Prostaglandins (PGs) can be synthesized through the activities of two cyclooxygenase (COX) isoforms. COX-1 is constitutively expressed in most tissues and its activity provides for the relative small amounts of PGs required for the mediation and modulation of normal physiological functions. In inflammatory conditions, ~~COX-2 is~~ rapidly induced by cytokines, growth factors and bacterial endotoxin, and its enzymatic activity accounts for the large amounts of PGs produced during inflammation. The currently used nonsteroidal anti-inflammatory drugs (NSAIDs) are nonselective inhibitors of both COX isoforms. Inhibition of COX-2 leads to the therapeutically desired inhibition of the synthesis of pro-inflammatory PGs, but at the same time produces side effects associated with inhibition of COX-1 and the consequent suppression of the production of PGs necessary for normal cellular functions. Selective inhibition of COX-2

expression explains, at least in part, the potent anti-inflammatory activity of corticosteroids. However, the systemic and ocular side effects of these steroids have greatly limited their use, especially their long-term use for the management of chronic inflammatory conditions. The current effort to develop highly selective nonsteroidal COX-2 inhibitors for the treatment of arthritis and other inflammatory diseases can also be expected to yield a new approach to the treatment of uveitis and other ocular inflammatory conditions. This new class of NSAIDs will provide anti-inflammatory and analgesic activity while circumventing the most serious side effects of the current available NSAIDs, resulting from their inhibition of the physiologically required COX-1 activity.

L137 ANSWER 15 OF 50 MEDLINE

ACCESSION NUMBER: 92195583 MEDLINE

DOCUMENT NUMBER: 92195583 PubMed ID: 1666177

TITLE: Treatment of experimental Pseudomonas keratitis with cyclo-oxygenase and lipoxxygenase inhibitors.

AUTHOR: Moreira H; McDonnell P J; Fasano A P; Silverman D L; Coates T D; Sevanian A

CORPORATE SOURCE: Doheny Eye Institute, Los Angeles, CA 90033.

CONTRACT NUMBER: EYO 3040 (NEI)

SOURCE: OPTHALMOLOGY, (1991-Nov)-98 (11) 1693-7.

JOURNAL CODE: 7802443. ISSN: 0161-6420.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199204

ENTRY DATE: Entered STN: 19920509

Last Updated on STN: 19960129

Entered Medline: 19920421

AB The role of metabolites of arachidonic acid in experimental Pseudomonas keratitis was studied using inhibitors of arachidonic acid metabolism. Nordihydroguaiaretic acid 1%, which inhibits predominantly the lipoxxygenase pathway, and flurbiprofen 0.03%, which inhibits predominantly the cyclo-oxygenase pathway were administered topically to rabbit eyes after intrastromal injection of Pseudomonas aeruginosa. Levels of the cyclo-oxygenase product prostaglandin E2 (PGE2) and the lipoxxygenase product leukotriene B4 (LTB4) were measured, and the number of ulcers that had progressed to descemetocoele formation by 24 hours was determined. Corneal ulceration was accelerated by flurbiprofen, but nordihydroguaiaretic acid limited the flurbiprofen-induced worsening. The use of flurbiprofen was associated with decreased levels of PGE2 and a relative increase in LTB4, a potent chemoattractant and activator of polymorphonuclear leukocytes. These results suggest that inhibition of the cyclo-oxygenase pathway may be contraindicated in Pseudomonas keratitis; inhibition of lipoxxygenase can prevent this worsening of the keratitis.

L137 ANSWER 16 OF 50 MEDLINE

ACCESSION NUMBER: 90147178 MEDLINE

DOCUMENT NUMBER: 90147178 PubMed ID: 2302115

TITLE: Reiter's Keratoconjunctivitis.

AUTHOR: Wiggins R E Jr; Steinkuller P G; Hamill M B

SOURCE: ARCHIVES OF OPTHALMOLOGY, (1990 Feb) 108 (2) 280-1.

JOURNAL CODE: 77065347. ISSN: 0003-9950.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199003

ENTRY DATE: Entered STN: 19900328

Last Updated on STN: 19900328

Entered Medline: 19900315

## L137 ANSWER 17 OF 50 MEDLINE

ACCESSION NUMBER: 89239600 MEDLINE  
DOCUMENT NUMBER: 89239600 PubMed ID: 3247209  
TITLE: [Piroxicam eyedrops in keratoconjunctivitis sicca. A new therapeutic perspective].  
Piroxicam collyre dans la keratoconjonctivite seche. Une nouvelle perspective therapeutique.  
AUTHOR: Bragliani G; Franco F; Marescotti A; Gaiba G  
SOURCE: OPHTALMOLOGIE, (1988 Oct) 2 (4) 359-62.  
Journal code: 8900549. ISSN: 0989-3105.  
PUB. COUNTRY: France  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: French  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 198906  
ENTRY DATE: Entered STN: 19900306  
Last Updated on STN: 19900306  
Entered Medline: 19890619

## L137 ANSWER 18 OF 50 MEDLINE

ACCESSION NUMBER: 87012857 MEDLINE  
DOCUMENT NUMBER: 87012857 PubMed ID: 3761968  
TITLE: [Local treatment with diclofenac-Na eyedrops in diseases of the anterior eye segment].  
Lokale Behandlung mit Diclofenac-Na-Augentropfen bei Erkrankungen der vorderen Augenabschnitte.  
AUTHOR: van Husen H  
SOURCE: KLINISCHE MONATSBLATTER FUR AUGENHEILKUNDE, (1986 Jun) 188 (6) 615-9.  
Journal code: 0014133. ISSN: 0023-2165.  
PUB. COUNTRY: GERMANY, WEST: Germany, Federal Republic of  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: German  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 198611  
ENTRY DATE: Entered STN: 19900302  
Last Updated on STN: 19900302  
Entered Medline: 19861104

AB The nonsteroid anti-inflammatory drug (NSAID) diclofenac-sodium, in the galenic form of an eye drop solution (0.1%), has been tested in an open clinical trial in the following indications: episcleritis (30 patients), limbal corneal ulcer (9 patients), hay fever conjunctivitis and/or conjunctivitis phlyctenulosa (11 patients). The result of this clinical trial has shown that diclofenac-sodium eye drop solution fulfills all the requirements of a well-tolerated and effective NSAID. The application of diclofenac-sodium eye drop solution (3-5 times daily) resulted in a clear-cut reduction in the use of eye drops containing steroids and its prominent analgesic effect was impressive. Although a slight, transient burning sensation was noticed by a few patients shortly after instillation, no local or systemic adverse reactions were observed.

## L137 ANSWER 19 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 2002-13547-DRUGU P  
TITLE: Nonsteroidal anti-inflammatory drugs prevent early diabetic retinopathy via TNF-alpha suppression.  
AUTHOR: Jousseaume A M; Poulaki V; Mitsiades N; Kirchhof B; Koizumi K; Doehman S; Adamis A P  
CORPORATE SOURCE: Harvard-Med.Sch.; Univ.Cologne  
LOCATION: Boston, Mass., USA; Cologne, Ger.  
SOURCE: FASEB J. (16, No. 3, 438-40, 2002) 3 Fig.  
CODEN: FAJOEC ISSN: 0892-6638  
AVAIL. OF DOC.: Retina Research Laboratory, Massachusetts Eye and Ear

Infirmery, Harvard Medical School, 324 Cambridge St., Boston, MA 02115, U.S.A. (A.P.A.). (e-mail: tony\_adamis@meei.harvard.edu).

LANGUAGE: English  
DOCUMENT TYPE: Journal  
FIELD AVAIL.: AB; LA; CT  
FILE SEGMENT: Literature

AB The effects of aspirin (AS), meloxicam (MX), and etanercept (ET) in the pathogenesis of diabetic retinopathy was investigated in a rat model of diabetic retinopathy. AS, MX, and ET reduced leukocyte adhesion, blood-retinal barrier breakdown, and TNF-alpha production. All 3 drugs also prevented the up-regulation of endothelial nitric oxide synthase (eNOS), ICAM-1 and the activation of nuclear factor kappa B (NF-kappa B). Only AS was able to down-regulate Erk kinase activation and leukocyte CD11a, CD11b, CD18 surface protein levels. Results indicate that these pharmacological agents had a beneficial effect in early experimental diabetic retinopathy and may hold promise for clinical efficacy in patients.

1137 ANSWER 20 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT  
ACCESSION NUMBER: 2002-08427 DRUGU S

TITLE: Unusual NSAID hypersensitivity.

AUTHOR: Fernandez Rivas M; Miranda T

LOCATION: Alcorcon, Esp.

SOURCE: Allergy (57, No. 2, 183-84, 2002) 3 Ref.

CODEN: LLRGDY ISSN: 0105-4538

AVAIL. OF DOC.: Fundacion Hospital Alcorcon, Unidad de Alergia, C/Budapest 1, 28922 Alcorcon, Spain.

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AB A case is reported of conjunctivitis induced by non-steroidal antiinflammatory drugs (NSAIDs: aspirin, metamizole, ibuprofen, diclofenac and dexametoprolen); no other such selective ocular reactions are thought to have been reported. Paracetamol and nimesulide were well tolerated. No protective effect was offered by premedication with disodium cromoglycate, sodium nedocromil, levocabastine and fluorometolone eye drops. In conclusion, this is an exceptional case of isolated left eye conjunctivitis after p.o. (and local) administration of NSAIDs, in which a local idiosyncratic reaction to inhibition of the cyclo-oxygenase pathway seems to be involved. The patient was advised to take paracetamol or nimesulide in future.

1137 ANSWER 21 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT  
ACCESSION NUMBER: 2001-31694 DRUGU T S

TITLE: Remitting seronegative symmetrical synovitis with pitting edema following intravesical bacillus Calmette-Guerin instillation.

AUTHOR: Mouly S; Berenbaum F; Kaplan G

LOCATION: Paris, Fr.

SOURCE: J.Rheumatol. (28, No. 7, 1699-701, 2001) 1 Tab. 15 Ref.

CODEN: JRHUA9 ISSN: 0315-162X

AVAIL. OF DOC.: General Clinical Research Center, Campus Box No. 7600, Room 3005 APCF, The University of North Carolina, Chapel Hill, NC 27599-7600, U.S.A. (e-mail: snouly@email.unc.edu).

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AB A case of remitting seronegative symmetrical synovitis with pitting edema following intravesical BCG in an HLA-B27 positive bladder carcinoma patient is described. The patient was admitted for polyarthritis after

receiving intravesical BCG. He received ketoprofen and morphine sulfate and showed immediate improvement. AST, ALT, serum gamma-glutamyltransferase, and alkaline phosphatases increased following ketoprofen treatment. Morphine was stopped and indometacin was started in place of ketoprofen. The patient still complained of moderate pain with synovitis in the ankles, knees, and joints and indometacin was replaced with meloxicam. This resulted in a complete resolution of joint pains, synovitis, knee effusions, and behavioral changes.

L137 ANSWER 22 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-47054 DRUGU P

TITLE: Pharmacological actions and therapeutic uses of cannabis and cannabinoids.

AUTHOR: Kumar R N; Chambers W A; Pertwee R G

CORPORATE SOURCE: Univ.Grampian

LOCATION: Aberdeen, U.K.

SOURCE: Anaesthesia (56, No. 11, 1059-68, 2001), 1 Tab. 79 Ref.

CODEN: ANASAB ISSN: 0003-2409

AVAIL. OF DOC.: Department of Anaesthesia, Grampian University Hospitals, Aberdeen AB25 2ZN, Scotland. (W.A.C.). (e-mail: alastair.chambers@arh.grampian.scot.nhs.uk).

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AB The pharmacological actions and therapeutic uses of cannabis and cannabinoids are reviewed. 2 Cannabis receptors (CB1 and CB2) bind the endogenous ligands anandamide, 2-arachidonoylglycerol and palmitoylethanol amide, the capsaicin analog olvanil and various synthetic compounds (WIN-55212, CP-55940, SR-144528 and SR-141716A) but some effects are mediated by non-receptor mechanisms. Tetrahydrocannabinol (THC) and other cannabinoids are rapidly absorbed and metabolised. Relaxant effects have led to use in spasticity (Nabilone), pain (levonatradol), emesis, anorexia, epilepsy, glaucoma, asthma and psychiatry, toxicity is low but sedation is common and tolerance can be induced.

L137 ANSWER 23 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-43235 DRUGU P

TITLE: Inhibition of COX in ocular tissues: An in vitro model to identify selective COX-2 inhibitors.

AUTHOR: Garcia Cabanes C; Palmero M; Bellot J L; Castillo M; Orts A

CORPORATE SOURCE: Univ.Alicante

LOCATION: Alicante, Esp.

SOURCE: J.Ocul.Pharmacol.Ther. (17, No. 1, 67-73, 2001) 3 Fig. 1 Tab. 28 Ref.

CODEN: JOPTF ISSN: 1080-7683

AVAIL. OF DOC.: Department of Interuniversity Optics, University of Alicante, Campus de San Vicente, E-03080 Alicante, Spain. (A.O.). (e-mail: alfredo.orts@ua.es).

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AB Incubation with diclofenac (Sigma-Chem.) or NS-398 (Calbiochem) resulted in inhibition of the lipopolysaccharide (LPS, Sigma-Chem.)-induced increase in PGE2 synthesis in both cultured bovine corneal endothelial cells (CEC) and retinal pigmentary epithelial (RPE) cells. Diclofenac seemed to be a COX-2 inhibitor because its IC50 value in RPE cells were similar to the IC50 value of NS-398. Whereas in CEC, NS-398 was several times more potent than diclofenac in inhibiting PGE2 synthesis induced by LPS. Piroxicam (Tocris) was th

weaker inhibitor on either type of cell. Findings suggest that this in vitro model could be used as a suitable assay system to determine the COX-2 selectivity of new NSAID during inflammatory events in ocular tissues.

L137 ANSWER 24 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 2000-46050 DRUGU P B E

TITLE: ~~Nepafenac~~, a unique nonsteroidal prodrug with potential utility in the treatment of trauma-induced ocular inflammation: I. assessment of anti-inflammatory efficacy.  
AUTHOR: Gamache D-A; Graff G; Brady M T; Spellman J M; Yanni J M  
CORPORATE SOURCE: Alcon  
LOCATION: Fort Worth, Tex., USA  
SOURCE: Inflammation (24, No. 4, 357-70, 2000) 6 Fig. 2 Tab. 20 Ref.  
CODEN: INELD4 ISSN: 0360-3997  
AVAIL. OF DOC.: Pharmaceutical Products Research, Alcon Research, Ltd., 6201 S. Freeway, Fort Worth, Texas, U.S.A.  
LANGUAGE: English  
DOCUMENT TYPE: Journal  
FIELD AVAIL.: AB; LA; CT  
FILE SEGMENT: Literature

AB The effect of nepafenac (NP) in trauma-induced ocular inflammation was investigated in-vivo in rabbits and in in-vitro experiments. Diclofenac (DC) and ~~anfenac~~ (AM) were used as reference compounds. New Zealand Albino rabbits (2-2.5 kg) received ocular NP, DC 50 ul 0.1% or saline followed by induction of trauma-induced inflammation 45 min later. In-vitro, NP and DC showed cyclooxygenase (COX)-1 inhibitory activity with IC50 of 64.3 and 0.12 uM, respectively. ~~AM-inhibited~~ COX-1 and COX-2 with IC50 of 0.25 and 0.15 uM, respectively. Ex-vivo, NP inhibited prostaglandin activity in the iris ciliary body (85-95%) and the retinoid/choroid (55%) for 6 and 4 hr, respectively. NP was longer acting than DC and not as effective. In-vivo, this was confirmed in the ocular inflammation model. Results show there should be further investigation for postoperative ocular inflammation. (No EX).

L137 ANSWER 25 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-22240 DRUGU P E

TITLE: Ocular inflammatory models.  
AUTHOR: Ogawa T  
CORPORATE SOURCE: Senju  
LOCATION: Osaka, Jap.  
SOURCE: Jpn J Pharmacol. (82, Suppl. 1, 198, 2000)  
CODEN: JJPAAZ ISSN: 0021-5198  
AVAIL. OF DOC.: International R & D Division, Senju Pharmaceutical Co., Osaka 541-0046, Japan.  
LANGUAGE: English  
DOCUMENT TYPE: Journal  
FIELD AVAIL.: AB; LA; CT  
FILE SEGMENT: Literature

AB The effects of topically applied cyclooxygenase (COX) inhibitors, bromfenac sodium (BF), betamethasone (BM), indometacin (IM) and nimesulide, on ocular inflammatory rat and rabbit models were investigated. The models showed that the COX isozyme involved in response was different in models and there were some models where prostaglandins (PGs) did not have any role in ocular signs. It was concluded that suitable models should be carefully selected to show the efficacy of COX inhibitors for clinical use. (conference paper: 73rd Annual Meeting of the Japanese Pharmacological Society, Yokohama, Japan, 2000.).

37 ANSWER 26 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT

ACCESSION NUMBER: 1999-22369 DRUGU P

Arterially perfused eye model ~~of uveitis~~

AUTHOR: Shiels I A; Sanderson S D; Taylor S M  
LOCATION: Brisbane, Austr.; Omaha, Neb., USA  
SOURCE: Aust.Vet.J. (77, No. 2, 100-104, 1999) 4 Fig. 22 Ref.  
CODEN: AUVJA2 ISSN: 0005-0423  
AVAIL. OF DOC.: Department of Physiology and Pharmacology, University of  
Queensland, St. Lucia, Queensland 4072, Australia.  
LANGUAGE: English  
DOCUMENT TYPE: Journal  
FIELD AVAIL.: AB; LA; CT  
FILE SEGMENT: Literature  
AB An in vitro model of uveitis based on an ex situ perfused eye was developed to evaluate the anti-inflammatory activity of new pharmacological products. Hydrogen peroxide reduced the intraocular pressure and perfusion flow rate in canine eyes. Flunixin meglumine, ketoprofen, indomethacin and pirfenidone (PFD) inhibited the effects of hydrogen peroxide on intraocular pressure, but not those on mediator-induced changes in perfusate flow. Uveitis involves inflammation of intraocular tissue. PFD is a novel antifibrotic drug currently being evaluated for activity in pulmonary fibrosis in humans. PFD may also show activity in other fibrosing diseases such as recurrent uveitis. The new model of uveitis should allow evaluation of anti-inflammatory activity without the need for experimental animals.

L137 ANSWER 27 OF 50 DRUGU COPYRIGHT 2002 THOMSON DERWENT  
ACCESSION NUMBER: 1998-15439 DRUGU T  
TITLE: Enzyme-inhibitors as drugs. (Part III).  
AUTHOR: Nuhn P  
CORPORATE SOURCE: Univ.Martin=Luther  
LOCATION: Halle, Ger.  
SOURCE: Pharm.Unserer Zeit (27, No. 1, 12-17, 1998) 33 Ref.  
CODEN: PHUZBI ISSN: 0048-3664  
AVAIL. OF DOC.: Fachbereich Pharmazie, Martin-Luther-Universitaet  
Halle-Wittenberg, Wolfgang-Langenbeck-Str. 4, 06120 Halle,  
Germany.  
LANGUAGE: German  
DOCUMENT TYPE: Journal  
FIELD AVAIL.: AB; LA; CT  
FILE SEGMENT: Literature  
AB The use of enzyme-inhibitors as drugs is reviewed with reference to inhibitors of the biosynthesis of mediators of inflammation, protease inhibitors, inhibitors of enzymes involved in carbohydrate and fat metabolism, and inhibitors of carbonic anhydrase. Protease inhibitors are used in treatment of coagulation disorders, hemorrhagic shock, septic shock, inflammatory diseases (pancreatitis, rheumatoid arthritis, acute respiratory syndrome, lung emphysema) and ulceration of the cornea. Inhibitors of carbohydrate metabolism can be used in combination with insulin to prevent accumulation of sorbitol and fructose. Inhibitors of carbonic anhydrase are used as diuretics and antiepileptics, and in treatment of glaucoma.

L137 ANSWER 28 OF 50 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2002162385 EMBASE  
TITLE: Can we prevent recurrences of herpes infections without antiviral drugs? The Weisenfeld Lecture.  
AUTHOR: Kaufman H.E.  
CORPORATE SOURCE: H.E. Kaufman, LSU Eye Center, 2020 Gravier Street, New Orleans, LA 70112, United States. hkaufm@lsuhsc.edu  
SOURCE: Investigative Ophthalmology and Visual Science, (2002) 43/5 (1325-1329).  
Refs: 37  
ISSN: 0146-0404 CODEN: IOVSDA  
COUNTRY: United States  
DOCUMENT TYPE: Journal; Conference Article

FILE SEGMENT: 012 Ophthalmology  
037 Drug Literature Index  
LANGUAGE: English

L137 ANSWER 29 OF 50 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2001362880 EMBASE  
TITLE: [Development of markets for over-the-counter drugs and food supplements in the USA 2000].  
ENTWICKLUNG DES MARKTES FUR OTC-ARZNEIMITTEL UND NAHRUNGSERGANZUNGSMITTEL IN DEN USA 2000.  
AUTHOR: Walluf-Blume D.  
CORPORATE SOURCE: Dr. D. Walluf-Blume, Referat Selbstmedikation, Bvb Pharmazeutischen Industrie e.V., Karlstr. 21, Frankfurt/Main, Germany  
SOURCE: Pharmazeutische Industrie, (2001) 63/9 (944-949).  
Refs: 27  
ISSN: 0031-711X CODEN: PHINAN  
COUNTRY: Germany  
DOCUMENT TYPE: Journal; Article  
FILE SEGMENT: 017 Public Health, Social Medicine and Epidemiology  
036 Health Policy, Economics and Management  
037 Drug Literature Index  
LANGUAGE: German

L137 ANSWER 30 OF 50 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2001212917 EMBASE  
TITLE: Cataract/IOL surgeries and postoperative pseudophakias - Related topics in the near future.  
AUTHOR: Miyake K.  
CORPORATE SOURCE: K. Miyake, Miyake Eye Hospital, 5-1070 Kami Higashi-Oosone-cho, Kita-ku Nagoya-shi 462-0823, Japan  
SOURCE: Japanese Journal of Clinical Ophthalmology, (2001) 55/5 (739-751).  
Refs: 15  
ISSN: 0370-5579 CODEN: RIGAA3  
COUNTRY: Japan  
DOCUMENT TYPE: Journal; Article  
FILE SEGMENT: 012 Ophthalmology  
037 Drug Literature Index  
038 Adverse Reactions Titles  
LANGUAGE: Japanese  
SUMMARY LANGUAGE: English; Japanese

AB Cataract/IOL surgery shows consistently good postoperative results, and is one of the successes of 20th century ophthalmology. With phacoemulsification, however, there is a regular incidence of intraoperative complications, postoperative complications such as aftercataracts, and problems such as the quality of postoperative vision. At the same time, since a huge number of patients undergo this procedure, surgical training issues remain. In relation to these problems, we herein discuss the preclinical evaluation of new techniques such as laser surgery to replace phacoemulsification, the possibility of selective COX-2 inhibiting nonsteroidal eyedrops in pseudophakic eyes, the mechanism of cystoid macular edema caused by anti-glaucoma eyedrops, and the application of new surgical observation systems using high-definition, high-quality 3D-TV for cataract/IOL surgery education.

L137 ANSWER 31 OF 50 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2000427683 EMBASE  
TITLE: Corneal foreign bodies.  
SOURCE: Practical Optometry, (2000) 1175 (191).  
ISSN: 1181-6058 CODEN: PROPFW  
COUNTRY: Canada  
DOCUMENT TYPE: Journal; General Review



FILE SEGMENT: 012 Ophthalmology  
027 Biophysics, Bioengineering and Medical  
Instrumentation  
037 Drug Literature Index  
LANGUAGE: English

L137 ANSWER 32 OF 50 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 2000327454 EMBASE  
TITLE: Fuchs' endothelial corneal dystrophy.  
AUTHOR: Melton R.; Thomas R.  
SOURCE: Practical Optometry, (2000) 11/4 (168-170).  
ISSN: 1181-6058 CODEN: PROPFW

COUNTRY: Canada  
DOCUMENT TYPE: Journal; Article  
FILE SEGMENT: 012 Ophthalmology  
037 Drug Literature Index  
LANGUAGE: English

L137 ANSWER 33 OF 50 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 1

ACCESSION NUMBER: 2002:293418 CAPLUS  
DOCUMENT NUMBER: 136:330549  
TITLE: Topical antibiotic composition for treatment of eye  
infection  
INVENTOR(S): Bandyopadhyay, Rebanta; Secreast, Pamela J.; Hawley,  
Leslie C.; McCurdy, Vincent E.; Tyle, Praveen;  
Bandyopadhyay, Paramita; Singh, Satish K.  
PATENT ASSIGNEE(S): Pharmacia & Upjohn Company, USA  
SOURCE: PCT Int. Appl., 41 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2002030395 | A1   | 20020418 | WO 2001-US31590 | 20011010 |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,  
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,  
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,  
PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,  
US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

|                        |    |          |                 |            |
|------------------------|----|----------|-----------------|------------|
| US 2002107238          | A1 | 20020808 | US 2001-974598  | 20011010   |
| PRIORITY APPLN. INFO.: |    |          | US 2000-239136P | P 20001010 |
|                        |    |          | US 2001-285340P | P 20010420 |

AB There is provided a pharmaceutical compn. suitable for topical  
administration to an eye; the compn. comprising as active agent one or  
more oxazolidinone antibacterial drugs, for example linezolid, in a concn.  
effective for treatment and/or prophylaxis of a gram-pos. bacterial  
infection of the eye, and one or more ophthalmically acceptable excipient  
ingredients that reduce rate of removal of the compn. from the eye by  
lacrimation such that the compn. has an effective residence time in the  
eye of about 2 to about 24 h. The compn. is, for example, an in situ  
gellable soln., suspension or soln./suspension. Formulations contg. a  
gelling or mucoadhesive agent (xanthan gum, HPMC, poloxamer 407, and  
polycarbophil) resulted in significant amts. of linezolid being retained  
in the exterior of treated eyes 1 h or more after application.

IT 162011-90-7, Rofecoxib 169590-42-5, Celecoxib  
181695-72-7, Valdecoxib 202409-33-4, MK-663

RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(topical antibiotic compn. for treatment of eye infection)

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L137 ANSWER 34 OF 50 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 2

ACCESSION NUMBER: 2002:71904 CAPLUS

DOCUMENT NUMBER: 136:112699

TITLE: Method of using cyclooxygenase 2 (  
COX-2) inhibitors in the  
treatment and prevention of ocular COX-  
2-mediated disorders

INVENTOR(S): Bandyopadhyay, Rebanta; Eveleth, David; Van Haarlem,  
Tom; Kararli, Tugrul T.; Singh, Satish K.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA

SOURCE: PCT Int. Appl., 103 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2002005848 | A2   | 20020124 | WO 2001-US14600 | 20010504 |
| WO 2002005848 | A3   | 20020704 |                 |          |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,  
HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,  
LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,  
RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,  
VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

US 2000-218101P P 20000713  
US 2001-279285P P 20010328

OTHER SOURCE(S): MARPAT 136:112699

AB. The invention provides methods for the treatment and prevention of ocular  
COX-2-mediated disorders using COX-2 inhibitors, e.g. celecoxib.

IT 329900-75-6, Cyclooxygenase 2

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(cyclooxygenase 2 inhibitors for  
treatment and prevention of ocular COX-2-mediated  
disorders)

IT 162011-90-7, Rofecoxib 169590-41-4, Deracoxib

169590-42-5, Celecoxib 181695-72-7,

Valdecoxib 202409-33-4, Etoricoxib

212126-32-4 266320-83-6

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)

(cyclooxygenase 2 inhibitors for  
treatment and prevention of ocular COX-2-mediated  
disorders)

L137 ANSWER 35 OF 50 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 3

ACCESSION NUMBER: 2002:71873 CAPLUS

DOCUMENT NUMBER: 136:123671

TITLE: Ophthalmic formulation of a selective  
cyclooxygenase-2 inhibitory  
drug

INVENTOR(S): Kararli, Tugrul T.; Bandyopadhyay, Rebanta; Singh,

PATENT ASSIGNEE(S): Satish K.; Hawley, Leslie C.  
SOURCE: Pharmacia & Upjohn Company, USA  
PCT Int. Appl., 71 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE  
-----  
WO 2002005815 A1 20020124 WO 2001-US22061 20010712  
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,  
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,  
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,  
RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,  
UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
US 2002035264 A1 20020321 US 2001-904098 20010712  
PRIORITY APPLN. INFO.: US 2000-218101P P 20000713  
US 2001-279285P P 20010328  
US 2001-294838P P 20010531  
US 2001-296388P P 20010606

OTHER SOURCE(S): MARPAT 136:123671

AB A pharmaceutical compn. suitable for topical administration to an eye contains a selective COX-2 inhibitor or nanoparticles of a drug of low water soly., at a concn. effective for the treatment and/or prophylaxis of a disorder in the eye, and 1 or more ophthalmically acceptable excipients that reduce rate of removal from the eye such that the compn. has an effective residence time of 2-24 h. Also provided is a method of treating and/or preventing a disorder in an eye, the method comprising administering to the eye a compn. of the invention. Thus, an ophthalmic nanoparticle suspension contained valdecoxib at 2.15 mg/g, 1.2% glycerin, 0.8% EDTA disodium salt, 4.0% Gelcarin GP-379NF, 0.21% SeaSpem PF and 0.82% Povidone.

IT 329900-75-6, Cyclooxygenase-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(inhibitor; ophthalmic formulation of  
cyclooxygenase-2 inhibitor pharmaceuticals)

IT 181695-72-7, Valdecoxib

RL: PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(ophthalmic formulation of cyclooxygenase-2  
inhibitor pharmaceuticals)

IT 162011-90-7, Rofecoxib 169590-41-4, Deracoxib

169590-42-5, Celecoxib 202409-33-4,

Etoricoxib 212126-32-4 266320-83-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(ophthalmic formulation of cyclooxygenase-2  
inhibitor pharmaceuticals)

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L137 ANSWER 36 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:521933 CAPLUS

DOCUMENT NUMBER: 137:108286

TITLE: Antibodies and fragments against epitopes present on  
cancer, metastatic or leukemia cells and platelets for  
diagnosis and therapy of tumor, metastasis, leukemia,  
autoimmune disease, and inflammation

INVENTOR(S): Lazarovits, Janette; Hagai, Yocheved; Plaksin, Daniel;  
Vogel, Tikva; Nimrod, Abraham; Mar-Haim, Hagit;  
Szanthon, Ester; Richter, Tamar; Amit, Boaz;  
Kooperman, Lena; Peretz, Tuvia; Levanon, Avigdor  
PATENT ASSIGNEE(S): Bio-Technology General Corp., USA  
SOURCE: PCT Int. Appl., 310 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

| PATENT NO.    | KIND   | DATE     | APPLICATION NO. | DATE     |
|---------------|--|----------|-----------------|----------|
| WO 2002053700 | A2   | 20020711 | WO 2001-US49442 | 20011231 |
| W:            | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |          |                 |          |
| RW:           | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |          |                 |          |

PRIORITY APPLN. INFO.: US 2000-258948P P 20001229  
US 2000-751181 A 20001229

AB The present invention provides epitopes present on cancer cells and important in physiological phenomena such as cell rolling, metastasis, and inflammation. Therapeutic and diagnostic methods and compounds using antibodies capable of binding to the epitopes are provided. The antibodies or fragments are capable of binding to, e.g. PSGL-1, fibrinogen, gamma. prime, GPIIb/IIIa, heparin, lumican, complement compd. 4 (C4), interalpha inhibitor and prothrombin. Methods and compounds according to the present invention can be used in diagnosis of and therapy for such diseases as cancer, including tumor growth and metastasis, leukemia, auto-immune disease, and inflammatory disease.

IT 162011-90-7, Rofecoxib 169590-42-5, Celecoxib

RL: DGN (Diagnostic use); THU (Therapeutic use); BIOL (Biological study);  
USES (Uses)

(antibodies and fragments against epitopes present on cancer, metastatic or leukemia cells and platelets for diagnosis and therapy of tumor, metastasis, leukemia, autoimmune disease, and inflammation)

L137 ANSWER 37 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:449662 CAPLUS

DOCUMENT NUMBER: 137:33310

TITLE: Preparation of aminopyrimidines as IKK inhibitors

INVENTOR(S): Kois, Adam; MacFarlane, Karen J.; Satoh, Yoshitaka;  
Bhagwat, Shripad S.; Parnes, Jason S.; Palanki, Moorthy S. S.; Erdman, Paul E.

PATENT ASSIGNEE(S): Signal Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 194 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND  | DATE     | APPLICATION NO. | DATE     |
|---------------|---|----------|-----------------|----------|
| WO 2002046171 | A2  | 20020613 | WO 2001-US46403 | 20011205 |
| W:            | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, |          |                 |          |

GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,  
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,  
PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA,  
UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,  
CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,  
BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2000-251816P P 20001206

OTHER SOURCE(S): MARPAT 137:33310

AB The title compds. [I; R1 = (un)substituted (hetero)aryl; R2 = H; R3 = H, alkyl; R4 = halo, OH, alkyl, alkoxy; R5, R6 = R8, (CH<sub>2</sub>)<sub>a</sub>COR<sub>9</sub>, (CH<sub>2</sub>)<sub>a</sub>CO<sub>2</sub>R<sub>9</sub>, etc.; or NR<sub>5</sub>R<sub>6</sub> = (un)substituted heterocycle; R8, R9 = H, alkyl, aryl, etc.; a = 0-4] having activity as inhibitors of IKK, particularly IKK-2, were prepd. E.g., a multi-step synthesis of I [R1 = 4-ClC<sub>6</sub>H<sub>4</sub>; R2-R6 = H] having an IC<sub>50</sub> of .ltoreq. 1 .mu.M in the IKK-2 enzyme assay, was given. Such compds. I have utility in the treatment of a wide range of conditions that are responsive to IKK inhibition. Thus, methods of treating such conditions are also disclosed, as are pharmaceutical compns. contg. one or more compds. of the above compds.

IT 162011-90-7, Rofecoxib 169590-42-5, Celecoxib

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(antiinflammatory agent; prepn. of anilinopyrimidines as IKK inhibitors)

L137 ANSWER 38 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:449661 CAPLUS

DOCUMENT NUMBER: 137:33309

TITLE: Preparation of anilinopyrimidines as JNK pathway inhibitors

INVENTOR(S): Kois, Adam; MacFarlane, Karen J.; Satoh, Yoshitaka;  
Bhagwat, Shripad S.; Parnes, Jason S.; Palanki,  
Moorthy S. S.; Erdman, Paul E.

PATENT ASSIGNEE(S): Signal Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 199 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND   | DATE     | APPLICATION NO. | DATE     |
|---------------|--|----------|-----------------|----------|
| WO 2002046170 | A2   | 20020613 | WO 2001-US46402 | 20011205 |
| W:            | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |          |                 |          |
| RW:           | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG   |          |                 |          |

PRIORITY APPLN. INFO.: US 2000-251904P P 20001206

OTHER SOURCE(S): MARPAT 137:33309

AB The title compds. [I; R1 = (un)substituted (hetero)aryl; R2 = H; R3 = H, alkyl; R4 = halo, OH, alkyl, alkoxy; R5, R6 = R8, (CH<sub>2</sub>)<sub>a</sub>COR<sub>9</sub>, (CH<sub>2</sub>)<sub>a</sub>CO<sub>2</sub>R<sub>9</sub>, etc.; or NR<sub>5</sub>R<sub>6</sub> = (un)substituted heterocycle; R8, R9 = H, alkyl, aryl, etc.; a = 0-4] having activity as inhibitors of the JNK pathway, were prepd. E.g., a multi-step synthesis of I [R1 = 4-ClC<sub>6</sub>H<sub>4</sub>; R2-R6 = H] having an IC<sub>50</sub> of .ltoreq. 10 .mu.M in the JNK2 assay, was given. Such compds. I have utility in the treatment of a wide range of conditions that are responsive to inhibition of the JNK pathway. Thus, methods of treating such conditions are also disclosed, as are pharmaceutical compns.

contg. one or more compds. of the above compds.

IT 162011-90-7, Rofecoxib 169590-42-5, Celecoxib  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(antiinflammatory agent; prepn. of anilinopyrimidines as JNK pathway inhibitors)

L137 ANSWER 39 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:540258 CAPLUS  
DOCUMENT NUMBER: 137:109267  
TITLE: Preparation of benzoxepinopyridines as HMG-CoA reductase inhibitors  
INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-chi; Sun, Chong-qing  
PATENT ASSIGNEE(S): USA  
SOURCE: U.S. Pat. Appl. Publ., 42 pp., Cont.-in-part of U.S. Ser. No. 875,155.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO.   | DATE     |
|------------------------|------|----------|-------------------|----------|
| US 2002094977          | A1   | 20020718 | US 2001-7407      | 20011204 |
| US 2002013334          | A1   | 20020131 | US 2001-875155    | 20010606 |
| PRIORITY APPLN. INFO.: |      |          | US 2000-211595P P | 20000615 |
|                        |      |          | US 2001-875155 A2 | 20010606 |

AB Title compds. I [X = O, S, SO, SO<sub>2</sub>, NR<sub>7</sub>; Z = HOCHCH<sub>2</sub>CH(OH)CH<sub>2</sub>CO<sub>2</sub>R<sub>3</sub>, 4-hydroxy-2-oxopyran-6-yl, etc.; n = 0, 1; R<sub>1</sub>, R<sub>2</sub> = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R<sub>3</sub> = H, alkyl, metal ion; R<sub>4</sub> = H, halo, CF<sub>3</sub>, etc.; R<sub>7</sub> = H, alkyl, aryl, alkanoyl, aroyl, alkoxycarbonyl, etc.; R<sub>9</sub>, R<sub>10</sub> = H, alkyl], were prepd. as HMG CoA reductase inhibitors active in inhibiting cholesterol biosynthesis, modulating blood serum lipids such as lowering LDL cholesterol and/or increasing HDL cholesterol, and treating hyperlipidemia, hypercholesterolemia, hypertriglyceridemia and atherosclerosis (no data). E.g., a multistep synthesis of II is reported.

IT 162011-90-7, Vioxx 169590-42-5, Celebrex  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(coadministered agents; prepn. of benzoxepinopyridines as HMG-CoA reductase inhibitors for the treatment of hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, and other disorders)

L137 ANSWER 40 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:392237 CAPLUS  
DOCUMENT NUMBER: 136:401651  
TITLE: Preparation of fused pyridine derivatives as HMG-CoA reductase inhibitors  
INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing  
PATENT ASSIGNEE(S): USA  
SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S. Ser. No. 875,218.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------|------|------|-----------------|------|
|------------|------|------|-----------------|------|

US 2002061901 A1 20020523 US 2001-8154 20011204  
US 2002028826 A1 20020307 US 2001-875218 20010606  
PRIORITY APPLN. INFO.: US 2000-211594P P 20000615  
US 2001-875218 A2 20010606

OTHER SOURCE(S): MARPAT 136:401651

AB The title compds. I and their pharmaceutically acceptable salts, esters, prodrug esters, and stereoisomers are claimed [wherein: Z = CH(OH)CH<sub>2</sub>CR<sub>7</sub>(OH)CH<sub>2</sub>CO<sub>2</sub>R<sub>3</sub> or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH<sub>2</sub>)<sub>x</sub> and/or (CH<sub>2</sub>)<sub>y</sub> together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R<sub>1</sub>, R<sub>2</sub> = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R<sub>3</sub> = H or lower alkyl; R<sub>4</sub> = H, halo, CF<sub>3</sub>, OH, alkyl, alkoxy, CO<sub>2</sub>H, (un)substituted NH<sub>2</sub>, cyano, (un)substituted CONH<sub>2</sub>, etc.; R<sub>7</sub> = H, alkyl]. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). I are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Prepn. of several compds. are described. For instance, a multistep synthesis of fused pyridine deriv. II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant)agonists of specific receptors, and as numerous named drugs.

IT 162011-90-7, Vioxx 169590-42-5,  
Celebrex

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(therapeutic compns. also contg.; prepn. of fused pyridine derivs. as HMG-CoA reductase inhibitors)

L137 ANSWER 41 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:833096 CAPLUS

DOCUMENT NUMBER: 135:352816

TITLE: Prevention of insulin-dependent diabetes, complications thereof, or allograft rejection by inhibition of cyclooxygenase-2 activity or inhibition of NF- $\kappa$ B activation

INVENTOR(S): Tabatabaie, Tahereh; Kotake, Yashige

PATENT ASSIGNEE(S): Oklahoma Medical Research Foundation, USA

SOURCE: PCT Int. Appl., 44 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.     | KIND | DATE     | APPLICATION NO. | DATE     |
|----------------|------|----------|-----------------|----------|
| WO 2001-085175 | A2   | 20011115 | WO 2001-US15174 | 20010510 |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2000-203572P P 20000511

AB Insulin-dependent diabetes mellitus (IDDM) is an autoimmune disease

believed to be caused by an inflammatory process in the pancreas leading to selective destruction of the .beta. cells. Inducible cyclooxygenase (COX-2) is expressed under inflammatory conditions and its product prostaglandin E2(PGE2) is an important inflammation mediator. Administration of the selective COX-2 inhibitor such as, e.g., NS-398 prevents the onset of diabetes in mice brought on by multiple low-doses of streptozotocin (STZ). Histol. observations indicated that STZ-mediated destruction of .beta. cells was prevented by NS-398 treatment. Delayed (day 3) administration of NS-398 was also protective in this model. These results demonstrate the crit. importance of COX-2 activity in autoimmune destruction of .beta. cells, and point to the fact that COX-2 inhibition should provide a preventive therapy against IDDM or other autoimmune problems, including allograft rejection. Inhibitors of NF-.kappa.B activation may also be used to prevent IDDM and allograft rejection.

L137 ANSWER 42 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:10616 CAPLUS

DOCUMENT NUMBER: 134:91125

TITLE: Pharmaceutical compositions containing aldose reductase inhibitors and selective cyclooxygenase-2 inhibitors

INVENTOR(S): Mylari, Banavara Lakshman

PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: Eur. Pat. Appl., 103 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE       |
|---|------|----------|-----------------|------------|
| EP 1064965  | A2   | 20010103 | EP 2000-305361  | 20000626   |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO |      |          |                 |            |
| JP 2001031569   | A2   | 20010206 | JP 2000-194053  | 20000628   |
| CA 2313063  | AA   | 20001230 | CA 2000-2313063 | 20000629   |
| BR 2000002957   | A    | 20010130 | BR 2000-2957    | 20000630   |
| PRIORITY APPLN. INFO.:  |      |          | US 1999-141695P | P 19990630 |

OTHER SOURCE(S): MARPAT 134:91125

AB: Pharmaceutical compns. contg. aldose reductase inhibitors, a prodrug thereof or a salts and selective cyclooxygenase-2 inhibitors, a prodrug thereof or salts thereof are disclosed. The compns. are used for the treatment of diabetic complications such as diabetic neuropathy, diabetic nephropathy, diabetic retinopathy and diabetic cardiomyopathy. Hard gelatin capsules contained active ingredients 0.25-100, starch 0.0-650, starch powder 0.0-50, and silicone fluid 350-cSt 0.15 mg/capsules.

L137 ANSWER 43 OF 50 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:10609 CAPLUS

DOCUMENT NUMBER: 134:76394

TITLE: Compositions containing aldose reductase inhibitors and selective cyclooxygenase inhibitors for the treatment of diabetic complications

INVENTOR(S): Mylari, Banavara Lakshman

PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: Eur. Pat. Appl., 11 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:



| PATENT NO.   | KIND | DATE     | APPLICATION NO. | DATE       |
|--|------|----------|-----------------|------------|
| EP 1064949   | A2   | 20010103 | EP 2000-305354  | 20000626   |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO  |      |          |                 |            |
| US 6426341   | B1   | 20020730 | US 2000-602793  | 20000623   |
| JP 2001031589  | A2   | 20010206 | JP 2000-194425  | 20000628   |
| CA 2313105   | AA   | 20001230 | CA 2000-2313105 | 20000629   |
| BR 2000002933  | A    | 20010130 | BR 2000-2933    | 20000630   |
| PRIORITY APPLN. INFO.:   |      |          | US 1999-141780P | P 19990630 |
| OTHER SOURCE(S): MARPAT 134:76394  |      |          |                 |            |
| AB Pharmaceutical compns. contain aldose reductase inhibitors such as zopolrestat and selective cyclooxygenase-2 inhibitors for the treatment of diabetic complications. |      |          |                 |            |
| L137 ANSWER 44 OF 50 CAPLUS COPYRIGHT 2002 ACS   |      |          |                 |            |
| ACCESSION NUMBER: 1999:529121 CAPLUS   |      |          |                 |            |
| DOCUMENT NUMBER: 131:157648  |      |          |                 |            |
| TITLE: Preparation of biarylacetic acid derivatives as COX-2 inhibitors  |      |          |                 |            |
| INVENTOR(S): Bayly, Christopher I.; Black, Cameron; Ouimet, Nathalie; Percival, David; Leger, Serge; Ouellet, Marc   |      |          |                 |            |
| PATENT ASSIGNEE(S): Merck Frosst Canada & Co., Can.  |      |          |                 |            |
| SOURCE: PCT Int. Appl., 66 pp.   |      |          |                 |            |
| CODEN: PIXXD2  |      |          |                 |            |
| DOCUMENT TYPE: Patent  |      |          |                 |            |
| LANGUAGE: English  |      |          |                 |            |
| FAMILY ACC. NUM. COUNT: 1  |      |          |                 |            |
| PATENT INFORMATION:  |      |          |                 |            |

| PATENT NO.   | KIND | DATE     | APPLICATION NO. | DATE       |
|--|------|----------|-----------------|------------|
| WO 99/1224   | A1   | 19990819 | WO 1999-CA120   | 19990211   |
| W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  |      |          |                 |            |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG   |      |          |                 |            |
| US 5994379   | A    | 19991130 | US 1999-246925  | 19990209   |
| CA 2318966   | AA   | 19990819 | CA 1999-2318966 | 19990211   |
| AU 9925065   | A1   | 19990830 | AU 1999-25065   | 19990211   |
| EP 1054857   | A1   | 20001129 | EP 1999-904652  | 19990211   |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI, LT, LV, FI, RO  |      |          |                 |            |
| JP 2002503647  | T2   | 20020205 | JP 2000-531421  | 19990211   |
| PRIORITY APPLN. INFO.:   |      |          | US 1998-74627P  | P 19980213 |
|  |      |          | WO 1999-CA120   | W 19990211 |
| OTHER SOURCE(S): MARPAT 131:157648   |      |          |                 |            |
| AB Title compds. [I; R = H, CH3; R2 = H, F; R3 = H, CH3; Y = C(OEt), C(OMe), N, CH, C=O; Z = C, N; A = H; B = OEt, SET, OPr, (E)-CH:CHCH3, CH3; A-B = NHC(CH3):CH; CHN(CH3)CH, OC(CH3):CH, SC(CH3):CH, NHC(CH3):N, N:C(CH3)O, N:C(CH3)S, OC(CH3):N, SC(CH3):N, CH2N(CH3)CH, CHC(CH3)N:CH; dotted bond = single, double in relation to Y, Z, A, B], pharmaceutically acceptable salts (sodium, potassium, calcium, magnesium), tautomer, and esters thereof are prep'd. and compns. which contain such compds. and methods of use the compds. are presented and tested as inhibitors of COX-2. Thus, the title compd. I (Y = C(OEt); Z = C; A = H; B = OEt; R = H; R2 = H; R3 = CH3; dotted bonds = double bonds) was prep'd. from 3,5-diethoxyphenol in 3 steps. |      |          |                 |            |
| REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS   |      |          |                 |            |

## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L137 ANSWER 45 OF 50  
ACCESSION NUMBER: 2002-269309 [31] WPIDS  
DOC. NO. CPI: C2002-079950  
TITLE: Molded article for administration to oral cavity to treat or prevent cyclooxygenase-2 mediated condition contains selective **cyclooxygenase-2 inhibitor**.  
DERWENT CLASS: B02 B03 B07  
INVENTOR(S): KARARLI, T T; KONTNY, M J; LE, T T  
PATENT ASSIGNEE(S): (KARA-I) KARARLI T T; (KONT-I) KONTNY M J; (LETT-I) LE T T; (PHAA) PHARMACIA CORP  
COUNTRY COUNT: 97  
PATENT INFORMATION:

| PATENT NO   | KIND | DATE     | WEEK      | LA | PG |
|---|------|----------|-----------|----|----|
| WO 2002015884   | A2   | 20020228 | (200231)* | EN | 38 |
| RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ<br>NL OA PT SD SE SL SZ TR TZ UG ZW   |      |          |           |    |    |
| W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK<br>DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR<br>KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PH PL PT RO<br>RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW |      |          |           |    |    |
| US 2002071857   | A1   | 20020613 | (200243)  |    |    |
| AU 2001085011   | A    | 20020304 | (200247)  |    |    |

## APPLICATION DETAILS:

| PATENT NO     | KIND           | APPLICATION     | DATE     |
|---------------|----------------|-----------------|----------|
| WO 2002015884 | A2             | WO 2001-US25762 | 20010817 |
| US 2002071857 | A1 Provisional | US 2000-226487P | 20000818 |
|               |                | US 2001-932537  | 20010817 |
| AU 2001085011 | A              | AU 2001-85011   | 20010817 |

## FILING DETAILS:

| PATENT NO     | KIND       | PATENT NO    |
|---------------|------------|--------------|
| AU 2001085011 | A Based on | WO 200215884 |

PRIORITY APPLN. INFO: US 2000-226487P 20000818; US 2001-932537  
20010817

AB WO 200215884 A UPAB: 20020516

NOVELTY - Molded article (A) comprises a selective **cyclooxygenase-2 inhibitor** (I) with a carrier system comprising at least one carbohydrate. The ingredients and their amounts in the molded article and a process for preparing the article are selected so that the article exhibits rapid disintegration in the oral cavity. The mouldable blend is prepared by a process step not requiring wet granulation.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for the preparation of (A) which comprises mixing the drug with the excipient carrier system and shaping a unit dose quantity of the blend in a mold.

ACTIVITY - Analgesic; Antiinflammatory; Cardiant; Vasotropic; Respiratory; Dermatological; Cytostatic; Nootropic; Neuroprotective; Antiallergic; Cerebroprotective; Antiarthritic; Antianemic; Antithyroid; Ophthalmological; Gynecological; Tocolytic.

No biological data is given.

MECHANISM OF ACTION - **Cyclooxygenase-2 inhibitor**.

USE - Used for administration to an oral cavity to treat or prevent a

cyclooxygenase-2 mediated condition such as disorders characterized by inflammation and pain and/or fever e.g. arthritis, rheumatoid arthritis, spondyloarthropathies, gouty arthritis, osteoarthritis, systemic lupus erythematosus and juvenile arthritis, asthma, bronchitis, menstrual cramps, preterm labor, tendinitis, bursitis, allergic neuritis, cytomegalovirus infectivity, apoptosis, lumbago, liver disease, hepatitis, psoriasis, eczema, acne, burns, dermatitis, sunburn, post-operative inflammation, inflammatory bowel disease, Crohn's disease, gastritis, ulcerative colitis, migraine, periarteritis nodosa, thyroiditis, aplastic anemia, type I diabetes, nephrotic syndrome, Behcet's syndrome, polymyositis, gingivitis, nephritis, hypersensitivity, myocardial ischemia, retinitis, scleritis, ~~epi~~scleritis, conjunctivitis, retinopathies, uveitis, ocular photophobia, pulmonary inflammation, cystic fibrosis, bone resorption, Alzheimers disease, neurodegeneration, stroke, trauma, dementia, allergic rhinitis, respiratory distress syndrome, pain, cancer, cardiovascular disorders such as atherosclerosis, arteriosclerosis, myocardial infarction, thrombosis and angiogenesis disorders.

ADVANTAGE - (I) Exhibits rapid disintegration in the oral cavity. The moldable blend is prepared by a process not requiring wet granulation, so that the overall process can be simplified, problems during granulation can be avoided, the article can have improved organoleptic qualities and exhibit improved resistance to breakage or attrition during handling, packaging and removal from a package, and greater flexibility can be obtained in the form of the molded article. They also have less harmful side effects than nonsteroidal antiinflammatory drugs and less gastrointestinal toxicity and irritation.

Dwg.0/0

L137 ANSWER 46 OF 50 WPIDS (C) 2002 THOMSON DERWENT  
ACCESSION NUMBER: 2002-139466 [18] WPIDS  
DOC. NO. CPI: C2002-042870  
TITLE: New 1-(benzothiazol-2-yl)pyrazole derivatives are selective cyclooxygenase-2 inhibitors for treating inflammatory diseases and pain.  
DERWENT CLASS: B02  
INVENTOR(S): AOTSUKA, T; ISHITANI, K; KATO, H; WAGATSUMA, N  
PATENT ASSIGNEE(S): (GREM) GRELAN PHARM CO LTD  
COUNTRY COUNT: 96  
PATENT INFORMATION:

| PATENT NO  | KIND | DATE     | WEEK      | LA | PG |
|--|------|----------|-----------|----|----|
| WO 2001087880  | A1   | 20011122 | (200218)* | JA | 32 |
| RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ<br>NL OA PT SD SE SL SZ TR TZ UG ZW  |      |          |           |    |    |
| W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK<br>DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR<br>KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU<br>SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW |      |          |           |    |    |
| AU 2001056705  | A    | 20011126 | (200222)  |    |    |

APPLICATION DETAILS:

| PATENT NO     | KIND | APPLICATION    | DATE     |
|---------------|------|----------------|----------|
| WO 2001087880 | A1   | WO 2001-JP3940 | 20010511 |
| AU 2001056705 | A    | AU 2001-56705  | 20010511 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|-----------|------|-----------|
|           |      |           |

AU 2001056705 A Based on WO 200187880

PRIORITY APPLN. INFO: JP 2000-141316 20000515

AB WO 200187880 A UPAB: 20020319

NOVELTY - 1-(Benzothiazol-2-yl)pyrazole derivatives (I) are new.

DETAILED DESCRIPTION - 1-(Benzothiazol-2-yl)pyrazole derivatives of formula (I) and their salts are new.

R1 = H, halo, lower alkyl or lower alkoxy;

R2 = lower haloalkyl or lower alkyl;

R3 = lower alkyl; and

n = 0-2.

ACTIVITY - Antiinflammatory; Analgesic; Antiarthritic; Antirheumatic; Osteopathic; Respiratory-Gen.; **Ophthalmological**.

1-(Benzothiazol-2-yl)-3-difluoromethyl-5-((4-methylsulfinyl)phenyl)pyrazole at 30 mg/kg orally suppressed 94% of adjuvant induced arthritis in rats compared to 64% for celecoxib at 30 mg/kg orally.

MECHANISM OF ACTION - Cyclooxygenase-Inhibitor-2.

USE - As selective cyclooxygenase-2 inhibitors ~~useful for treating and preventing inflammatory diseases~~ and pain (claimed) including rheumatoid arthritis, osteoarthritis, neuralgia, bronchitis, conjunctivitis, prostatic inflammation or gingivitis.

ADVANTAGE - Are selective and have reduced side effects such as gastric mucosa disorders.

Dwg.0/0

L137 ANSWER 47 OF 50 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-625725 [72] WPIDS

DOC. NO. CPI: C2001-186385

TITLE: Antagonizing the binding of an integrin to its ligand  
useful for the treatment of angiogenesis comprises  
administration of an ADAM-disintegrin domain polypeptide.

DERWENT CLASS: B04 B05 D16

INVENTOR(S): BLACK, R A; CERRETTI, D P; FANSLOW, W C; POINDEXTER, K M

PATENT ASSIGNEE(S): (IMMV) IMMUNEX CORP; (BLAC-I) BLACK R A; (CERR-I)  
CERRETTI D P; (FANS-I) FANSLOW W C; (POIN-I) POINDEXTER K M

COUNTRY COUNT: 94

PATENT INFORMATION:

| PATENT NO | KIND | DATE | WEEK | LA | PG |
|-----------|------|------|------|----|----|
|-----------|------|------|------|----|----|

|               |    |          |           |    |    |
|---------------|----|----------|-----------|----|----|
| WO 2001062905 | A2 | 20010830 | (200172)* | EN | 66 |
|---------------|----|----------|-----------|----|----|

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ  
NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM  
DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC  
LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE  
SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001047219 A 20010903 (200202)

US 2002042368 A1 20020411 (200227)

APPLICATION DETAILS:

| PATENT NO     | KIND           | APPLICATION     | DATE     |
|---------------|----------------|-----------------|----------|
| WO 2001062905 | A2             | WO 2001-US5701  | 20010223 |
| AU 2001047219 | A              | AU 2001-47219   | 20010223 |
| US 2002042368 | A1 Provisional | US 2000-184865P | 20000225 |
|               |                | US 2001-792200  | 20010223 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|-----------|------|-----------|
|-----------|------|-----------|

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AU 2001047219 A Based on

WO 200162905

PRIORITY APPLN. INFO: US 2000-184865P 20000225; US 2001-792200  
20010223

AB WO 200162905 A UPAB: 20011206

NOVELTY - Antagonizing the binding of an integrin to its ligand or inhibiting angiogenesis in a mammal in need of it comprising the administration of an ADAM-disintegrin domain polypeptide (preferably except an RGD sequence), is new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for identifying a compound that modulates integrin biological activity, interaction between an integrin and the ADAM disintegrin domain; or inhibits endothelial cell migration and/or angiogenesis involving combining a test compound with endothelial cells and with the ADAM-disintegrin domain polypeptide (I) that binds to the integrin or endothelial cells; and determining whether the test compound alters the binding of (I) to the integrin or the endothelial cells.

ACTIVITY - Integrin binding activity; antiinflammatory; osteopathic; vasotropic; thrombolytic.

MECHANISM OF ACTION - Endothelial cell migration inhibitor; angiogenesis inhibitor; integrin antagonist; neovascularization inhibitor. A planer endothelial cell migration assay was used to quantitate the inhibition of angiogenesis by ADAM-disintegrin-Fc-polypeptides in vitro. Primary human renal microvascular endothelial cells, (HRMEC) were isolated, cultured and used at the third passage after thawing. Replicate circular lesions wounds were generated in confluent HRMEC monolayers using a silicon-tipped drill press. At the time of wounding the medium was supplemented with 20 ng/ml phorbol-12-myristate-13 acetate (PMA) and/or a range of concentration of ADAM-disintegrin-Fc-polypeptide; ADAM-20 and -23 dis-Fc polypeptides showed the greatest inhibition of both EGF and PMA induced endothelial migration of 15  $\mu$ g/ml. While HuIgG (control) did not inhibit EGF or PMA induced endothelial cell.

USE - For treatment of ocular disorders, malignant and metastatic conditions, inflammatory diseases, osteoporosis, and other conditions mediated by accelerated bone resorption, restenosis, inappropriate platelet activation, recruitment or aggregation, thrombosis or a condition requiring aggregation, thrombosis or a condition requiring tissue repair or wound healing, angiogenesis, ocular neovascularization or solid tumor (all claimed); for the treatment of diabetic retinopathy, retinopathy or prematurity, neovascular glaucoma, retinoblastoma, retrolental fibroplasias, rubeosis, uveitis, macular degeneration, and corneal graft neovascularization, inflammatory diseases, ocular tumors, diseases associated with choroidal or iris neovascularization, arthritis, rheumatism, inflammatory bowel disease, psoriasis, coronary artery disease or injury, myocardial infarction or injury following myocardial infarction, stroke, unstable angina, atherosclerosis, arteriosclerosis, preeclampsia, embolism, platelet-associated ischemic disorders including lung ischemia, coronary ischemia, cerebral ischemia, restenosis following percutaneous coronary intervention including angioplasty, atherectomy, stent placement, and bypass surgery, thrombotic disorders including coronary artery thrombosis, cerebral artery thrombosis, intracardiac thrombosis, peripheral artery thrombosis, venous thrombosis, thrombosis and coagulopathies associated with exposure to a foreign or injured tissue surface and reocclusion following thrombosis, deep venous thrombosis, pulmonary embolism, transient ischemic attacks, and another conditions where vascular occlusion is a common underlying feature, in individuals at high risk for thrombus formation of reformation, advanced coronary artery disease, or for occlusion, reocclusion, stenosis and/or restenosis of blood vessels or stroke benign tumors and preneoplastic conditions, myocardial angiogenesis, hemophilic joints, scleroderma, vascular adhesions, asthma and allergy, eczema and dermatitis, graft versus host disease, sepsis, adult respiratory distress syndrome, telangiectasia, and

wound granulation. The method are used in combination with angioplasty procedures, such as balloon angioplasty, laser angioplasty, coronary atherectomy or similar techniques, carotid endarterectomy, anastomosis of vascular grafts, surgery having a high risk of thrombus formation (i.e. coronary bypass surgery, insertion of a prosthetic valve or vessel and the like), atherectomy, stent placement, placement of a chronic cardiovascular device such as an in-dwelling catheter or prosthetic valve or vessel, organ transplantation, or bypass surgery.  
Dwg.0/0

L137 ANSWER 48 OF 50 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-432701 [46] WPIDS

CROSS REFERENCE: 2001-417847 [44]; 2001-451589 [48]; 2001-451615 [48];  
2001-457263 [49]; 2001-475686 [51]; 2001-502457 [55];  
2002-089529 [12]

DOC. NO. CPI: C2001-130904

TITLE: Amorphous celecoxib, which has improved bioavailability and dissolution properties, is useful for treatment of disorders mediated by cyclooxygenase-2, e.g. inflammation or pain.

DERWENT CLASS: B03

INVENTOR(S): HAGEMAN, M J; HE, X; KARARLI, T T; MACKIN, L A; MIYAKE, P J; ROHRS, B R; STEFANSKI, K J

PATENT ASSIGNEE(S): (PHAA) PHARMACIA CORP; (PHAR-N) PHARM CORP

COUNTRY COUNT: 95

PATENT INFORMATION:

| PATENT NO | KIND | DATE | WEEK | LA | PG |
|-----------|------|------|------|----|----|
|-----------|------|------|------|----|----|

|               |    |          |           |    |    |
|---------------|----|----------|-----------|----|----|
| WO 2001042221 | A1 | 20010614 | (200146)* | EN | 43 |
|---------------|----|----------|-----------|----|----|

RW: AT BE CH CY DE DK EA ES FI FR GE GH GM GR IE IT KE LS LU MC MW MZ  
NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM  
DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC  
LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE  
SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

|               |   |          |          |  |  |
|---------------|---|----------|----------|--|--|
| AU 2001019311 | A | 20010618 | (200161) |  |  |
|---------------|---|----------|----------|--|--|

|            |    |          |          |    |  |
|------------|----|----------|----------|----|--|
| EP 1150959 | A1 | 20011107 | (200168) | EN |  |
|------------|----|----------|----------|----|--|

R: AL AT BE CH CY DE DK ES FI FR GE GR IE IT LI LT LU LV MC MK NL PT  
RO SE SI TR

|               |   |          |          |  |  |
|---------------|---|----------|----------|--|--|
| NO 2001003855 | A | 20011005 | (200171) |  |  |
|---------------|---|----------|----------|--|--|

|               |    |          |          |  |  |
|---------------|----|----------|----------|--|--|
| CZ 2001003210 | A3 | 20020313 | (200223) |  |  |
|---------------|----|----------|----------|--|--|

|               |   |          |          |  |  |
|---------------|---|----------|----------|--|--|
| BR 2000008058 | A | 20020326 | (200229) |  |  |
|---------------|---|----------|----------|--|--|

|               |    |          |          |  |  |
|---------------|----|----------|----------|--|--|
| SK 2001001268 | A3 | 20020702 | (200253) |  |  |
|---------------|----|----------|----------|--|--|

APPLICATION DETAILS:

| PATENT NO     | KIND | APPLICATION     | DATE     |
|---------------|------|-----------------|----------|
| WO 2001042221 | A1   | WO 2000-US32435 | 20001206 |
| AU 2001019311 | A    | AU 2001-19311   | 20001206 |
| EP 1150959    | A1   | EP 2000-982255  | 20001206 |
|               |      | WO 2000-US32435 | 20001206 |
| NO 2001003855 | A    | WO 2000-US32435 | 20001206 |
|               |      | NO 2001-3855    | 20010808 |
| CZ 2001003210 | A3   | WO 2000-US32435 | 20001206 |
|               |      | CZ 2001-3210    | 20001206 |
| BR 2000008058 | A    | BR 2000-8058    | 20001206 |
|               |      | WO 2000-US32435 | 20001206 |
| SK 2001001268 | A3   | WO 2000-US32435 | 20001206 |
|               |      | SK 2001-1268    | 20001206 |

FILING DETAILS:

| PATENT NO     | KIND        | PATENT NO    |
|---------------|-------------|--------------|
| AU 2001019311 | A Based on  | WO 200142221 |
| EP 1150959    | A1 Based on | WO 200142221 |
| CZ 2001003210 | A3 Based on | WO 200142221 |
| BR 2000008058 | A Based on  | WO 200142221 |
| SK 2001001268 | A3 Based on | WO 200142221 |

PRIORITY APPLN. INFO: US 2000-169856 20001201; US 1999-169856P  
19991208; US 2000-730663 20001201

AB WO 200142221 A UPAB: 20020820

NOVELTY - Amorphous celecoxib is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for:

(A) amorphous celecoxib;

(B) celecoxib drug substance in which at least a portion of the celecoxib present is amorphous;

(C) celecoxib-crystallization inhibitor composite comprising particles of a material as described in (A) or (B) in intimate association with one or more crystallization inhibitors which reduces transformation of amorphous celecoxib to crystalline celecoxib; and

(D) composition comprising

(i) a material as described in (A), (B) or (C), in an amount which provides a total celecoxib dosage of 10-1,000 mg, and

(ii) one or more excipients.

ACTIVITY - Analgesic; Antipyretic; Antiinflammatory; Neuroprotective; Antiasthmatic; Antiseborrheic; Hypotensive; Cardioprotective; Cytostatic; Hepatotrophic; Dermatological; Ophthalmological; Antiallergic; Vulnerary; Gynecological; Osteopathic.

MECHANISM OF ACTION - Cyclooxygenase-2 inhibitor.

USE - ~~Celecoxib~~ is useful in treatment of disorders mediated by cyclooxygenase-2, including disorders characterized by inflammation, pain or fever. It can be used in treatment of, e.g. arthritis, asthma, bronchitis, menstrual cramps, pre-term labor, tendinitis, bursitis, neuritis, cytomegalovirus infectivity, lumbago, liver diseases, eczema, acne, burns, glaucoma, dermatitis, gastrointestinal conditions, ophthalmic diseases, pulmonary inflammation, nervous system disorders, allergic rhinitis, respiratory distress syndrome, endotoxin shock syndrome, pain, inflammation-related cardiovascular disorders, angiogenesis-related disorders, or cancer. It can be used to decrease bone loss and to inhibit prostanoid-induced smooth muscle contraction, e.g. for treatment of dysmenorrhea.

ADVANTAGE - The amorphous exhibits enhanced bioavailability and improved dissolution properties, relative to crystalline celecoxib. It can be storage stable, particularly when combined with a crystallization inhibitor.

Dwg.0/5

L137 ANSWER 49 OF 50 WPIDS (C) 2002 THOMSON DERWENT

ACCESSION NUMBER: 2001-451589 [48] WPIDS

CROSS REFERENCE: 2001-417847 [44]; 2001-432701 [46]; 2001-451615 [48];  
2001-457263 [49]; 2001-475686 [51]; 2001-502457 [55];  
2002-089529 [12]; 2002-225919 [28]

DOC. NO. CPI: C2001-136348

TITLE: New oral valdecoxib compositions which have good bioavailability and a rapid onset of activity, are useful in treatment of disorders mediated by cyclooxygenase-2, e.g., arthritis.

DERWENT CLASS: B03 B07

INVENTOR(S): DESAI, S; KARARLI, T T; KONTNY, M J; NADKARNI, S

PATENT ASSIGNEE(S): (PHAA) PHARMACIA CORP; (PHAR-N) PHARM CORP

COUNTRY COUNT: 95

PATENT INFORMATION:

| PATENT NO  | KIND | DATE     | WEEK      | LA | PG |
|--|------|----------|-----------|----|----|
| WO 2001041762  | A2   | 20010614 | (200148)* | EN | 30 |
| RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ<br>NL OA PT SD SE SL SZ TR TZ UG ZW  |      |          |           |    |    |
| W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM<br>DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC<br>LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE<br>SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW |      |          |           |    |    |
| AU 2001019310  | A    | 20010618 | (200161)  |    |    |
| EP 1165072   | A2   | 20020102 | (200209)  | EN |    |
| R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL RO<br>SI   |      |          |           |    |    |
| SK 2001001269  | A3   | 20020404 | (200232)  |    |    |
| CZ 2001003163  | A3   | 20020612 | (200251)  |    |    |

## APPLICATION DETAILS:

| PATENT NO     | KIND | APPLICATION     | DATE     |
|---------------|------|-----------------|----------|
| WO 2001041762 | A2   | WO 2000-US32433 | 20001206 |
| AU 2001019310 | A    | AU 2001-19310   | 20001206 |
| EP 1165072    | A2   | EP 2000-982254  | 20001206 |
|               |      | WO 2000-US32433 | 20001206 |
| SK 2001001269 | A3   | WO 2000-US32433 | 20001206 |
|               |      | SK 2001-1269    | 20001206 |
| CZ 2001003163 | A3   | WO 2000-US32433 | 20001206 |
|               |      | CZ 2001-3163    | 20001206 |

## FILING DETAILS:

| PATENT NO     | KIND        | PATENT NO    |
|---------------|-------------|--------------|
| AU 2001019310 | A Based on  | WO 200141762 |
| EP 1165072    | A2 Based on | WO 200141762 |
| SK 2001001269 | A3 Based on | WO 200141762 |
| CZ 2001003163 | A3 Based on | WO 200141762 |

PRIORITY APPLN. INFO: US 2000-202269P 20000505; US 1999-169856P  
19991208; US 2000-181635P 20000210

AB: WO 200141762 A UPAB: 20020812

NOVELTY - Oral valdecoxib compositions which contain 1-100 mg of valdecoxib per dose and which meet specified pharmacokinetic requirements are new.

DETAILED DESCRIPTION - Pharmaceutical composition comprises:

- (i) 1-100 mg of valdecoxib per dose; and
- (ii) one or more excipients.

Upon oral administration of a single dose to a fasting subject, the time course of blood serum concentration is at least one of the following:

(a) a time to reach a threshold concentration for therapeutic effect not greater than 0.5 hours after administration;

(b) a time to reach maximum concentration (Tmax) not greater than 3 hours after administration; and/or

(c) a maximum concentration (Cmax) not less than 100 ng/ml.

ACTIVITY - Analgesic; antipyretic; antiinflammatory; neuroprotective; antiasthmatic; antiacne; hypotensive; cardiant; cytostatic; antiviral.

MECHANISM OF ACTION - Cyclooxygenase-2 inhibitor.

USE - Valdecoxib is useful in treatment of disorders mediated by cyclooxygenase-2, including disorders characterized by inflammation, pain or fever. It can be used in treatment of, e.g., arthritis, asthma, bronchitis, menstrual cramps, preterm labor, tendinitis, bursitis, neuritis, cytomegalovirus infectivity, lumbago,



liver diseases, eczema, acne, burns, glaucoma, dermatitis, gastrointestinal conditions, **ophthalmic** diseases, pulmonary inflammation, nervous system disorders, allergic rhinitis, respiratory distress syndrome, endotoxin shock syndrome, pain, inflammation-related cardiovascular disorders, angiogenesis-related disorders, or cancer. It can be used to decrease bone loss and to inhibit prostanoid-induced smooth muscle contraction, e.g., for treatment of dysmenorrhea.

ADVANTAGE - The composition has good bioavailability characteristics and has a rapid onset of activity.  
Dwg.0/4

L137 ANSWER 50 OF 50 WPIDS (C) 2002 THOMSON DERWENT  
ACCESSION NUMBER: 2000-181828 [16] WPIDS  
DOC. NO. CPI: C2000-056752  
TITLE: **Use of pyrazolylphenylsulfonfyl cyclooxygenase-2 inhibitors** for the treatment of angiogenesis mediated disorders e.g. metastasis, corneal graft rejection, gastric ulcer and ocular neovascularization.  
DERWENT CLASS: B03  
INVENTOR(S): MASFERRER, J; RAZ, A  
PATENT ASSIGNEE(S): (SEAR) SEARLE & CO G D  
COUNTRY COUNT: 1  
PATENT INFORMATION:

| PATENT NO  | KIND | DATE     | WEEK      | LA | PG |
|------------|------|----------|-----------|----|----|
| US 6025353 | A    | 20000215 | (200016)* |    | 15 |

## APPLICATION DETAILS:

| PATENT NO  | KIND | APPLICATION    | DATE     |
|------------|------|----------------|----------|
| US 6025353 | A    | US 1997-974201 | 19971119 |

PRIORITY APPLN. INFO: US 1997-974201 19971119

AB US 6025353 A UPAB: 20000330

NOVELTY - Treatment of angiogenesis-mediated disorders selected from metastasis, corneal graft rejection, ocular neovascularization, retinal neovascularization, diabetic retinopathy, retrolental fibroplasia, neovascular glaucoma, gastric ulcer, infantile hemangiomas, angiofibroma of the nasopharynx, avascular necrosis of bone, and endometriosis comprises administration of pyrazolylphenylsulfonfyl **cyclooxygenase-2 inhibitors**

DETAILED DESCRIPTION - Treatment of angiogenesis-mediated disorders selected from metastasis, corneal graft rejection, ocular neovascularization, retinal neovascularization, diabetic retinopathy, retrolental fibroplasia, neovascular glaucoma, gastric ulcer, infantile hemangiomas, angiofibroma of the nasopharynx, avascular necrosis of bone, and endometriosis comprises administration of pyrazolylphenylsulfonfyl **cyclooxygenase-2 inhibitors** of formula (I).

A = pyrazolyl;

R1 = heterocyclyl, cycloalkyl, cycloalkenyl, or aryl (optionally substituted);

R2 = CH3 or NH2; and

R3 = H, halo, alkyl, alkenyl, alkynyl, oxo, CN, carboxyl, cyanoalkyl, heterocycloxy, alkoxy, alkylthio, alkylcarbonyl, cycloalkyl, aryl, haloalkyl, heterocyclyl, cycloalkenyl, aralkyl, heterocycloalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkyloxyalkyl, alkoxyaralkoxyalkyl, alkoxyaralkoxyalkyl, aminocarbonyl, aminocarbonylalkyl,

alkylaminocarbonyl, N-arylaminoalkyl, N-alkyl-N-aryl-aminocarbonyl, alkylaminocarbonylalkyl, carboxyalkyl, alkylamino, N-arylamino, N-aralkylamino, N-alkyl-N-aralkylamino, N-alkyl-N-arylamino, aminoalkyl, alkylaminoalkyl, N-arylaminoalkyl, N-aralkylaminoalkyl, N-alkyl-N-arylaminoalkyl, N-alkyl-N-arylaminoalkyl, aryloxy, aralkoxy, arylthio, aralkylthio, alkylsulfonyl, alkylsulfonyl, aminosulfonyl, alkylaminosulfonyl, N-arylaminosulfonyl, arylsulfonyl, or N-alkyl-N-arylaminosulfonyl.

An INDEPENDENT CLAIM is also included for the treatment of angiogenesis-mediated disorders as above comprising administration of aminosulfonylphenylpyrazole derivatives of formula (II).

R4 = H, alkyl, haloalkyl, alkoxycarbonyl, CN, cyanoalkyl, carboxyl, aminocarbonyl, alkylaminocarbonyl, cycloalkylaminocarbonyl, arylaminocarbonyl, carboxyalkylaminocarbonyl, carboxyalkyl, aralkylcarbonylalkylaminocarbonyl, aminocarbonylalkyl, alkoxycarbonylcycloalkenyl, or hydroxyalkyl;

R5 = H, alkyl, CN, hydroxyalkyl, cycloalkyl, alkylsulfonyl, or halo;

R6 = aralkenyl, aryl, cycloalkyl, cycloalkenyl, or heterocyclyl (all optionally substituted by one or more Q1); and

Q1 = halo, alkylthio, alkylsulfonyl, CN, NO2, haloalkyl, alkyl, OH, alkenyl, hydroxyalkyl, carboxyl, cycloalkyl, alkylamino, dialkylamino, alkoxycarbonyl, aminocarbonyl, alkoxy, haloalkoxy, sulfamyl, heterocyclyl or amino.

ACTIVITY - Cytostatic; ophthalmological; immunosuppressive; antidiabetic; antiulcer; osteopathic; gynecological.

Effects of (I) on angiogenesis in vivo were evaluated using the mouse corneal neovascularization assay according to Muthukkaupiah et al., J. Natl. Cancer Inst., 69, 699-708 (1982). 4-(5-(4-Chlorophenyl)-3-difluoromethyl-pyrazol-1-yl)-benzenesulfonamide (Ia) inhibited fibroblast growth factor-induced angiogenesis in mice at a dose of 6 mg/kg/day.

MECHANISM OF ACTION - Cyclooxygenase-2 inhibitor; antimetastatic.

USE - The method is used for the treatment of angiogenesis-mediated disorders selected from metastasis, corneal graft rejection, ocular neovascularization, retinal neovascularization, diabetic retinopathy, retrolental fibroplasia, neovascular glaucoma, gastric ulcer, infantile hemangiomas, angiofibroma of the nasopharynx, avascular necrosis of bone, and endometriosis (claimed).

ADVANTAGE - (I) and (II) are selective cyclooxygenase-2 inhibitors.

Dwg. 0/0

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